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(71) Applicant (*for AE, AG, AL, AM, AT, AU, AZ, BA, BB, BE, BF, BG, BJ, BR, BW, BY, BZ, CA, CF, CG, CH, CI, CM, CN, CO, CR, CU, CY, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, FR, GA, GB, GD, GE, GH, GM, GN, GQ, GR, GW, HR, HU, ID, IE, IL, IN, IS, IT, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MC, MD, MK, ML, MN, MR, MW, MX, MZ, NA, NE, NG, NI, NL, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG only*): **ASTRAZENECA AB** [SE/SE]; SE-151 85 Södertälje (SE).

(71) Applicant (*for MG only*): **ASTRAZENECA UK LIMITED** [GB/GB]; 15 Stanhope Gate, London Greater London W1K 1LN (GB).

(72) Inventors; and

(75) Inventors/Applicants (*for US only*): **HUDSON, Kevin** [GB/GB]; AstraZeneca R & D Alderley, Alderley Park, Macclesfield Cheshire SK10 4TG (GB). **SOUTH, Marie**,

Caroline [GB/GB]; AstraZeneca R & D Alderley, Alderley Park, Macclesfield Cheshire SK10 4TG (GB). **MARSHALL, Gayle** [GB/GB]; AstraZeneca R & D Alderley, Alderley Park, Macclesfield Cheshire SK10 4TG (GB). **SAM, Mehran** [CA/US]; AstraZeneca R & D Boston, 35 Gatehouse Drive, Waltham, MA 02451 (US).

(74) Agent: **GLOBAL INTELLECTUAL PROPERTY**; AstraZeneca AB, SE-151 85 Södertälje (SE).

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(54) Title: METHOD

(57) Abstract: The invention relates to a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or 2 as defined herein whereby to predict an increased likelihood of response to the erbB receptor drug. Preferred genes include any one of NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2, VAMP4, DAPK1, DAPK2, MLLT3, TNNC1, KIAA0931, ACOX2, EMP1, SLC20A1, SPRY2 or PGM1.

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METHOD

The present invention relates to sensitivity of tumours to therapeutic agents which can be predicted from the gene expression profile of the tumour and hence that the suitability of cancer patients for treatment with such therapeutic agents can be determined by measuring the relative expression levels of particular genes in tumour tissue.

The phosphorylation of proteins on tyrosine residues is a key element of signal transduction within cells. Enzymes capable of catalysing such reactions are termed tyrosine kinases. A number of these enzymes exist as integral components of transmembrane receptor molecules and are classified as receptor tyrosine kinases (RTKs). There are several members of this family of RTKs, class I of which includes the erbB family, e.g. epidermal growth factor receptor (EGFR), erbB2, erbB3 and erbB4. Binding of a variety of ligands to the external domain activates the EGFR tyrosine kinase domain. Activation causes EGFR itself and a number of cellular substrates to become phosphorylated on tyrosine residues. These phosphorylation reactions are a major component of growth factor induced proliferation of cells.

The erbB family of receptor tyrosine kinases are known to be frequently involved in driving the proliferation and survival of tumour cells (reviewed in Olayioye *et al.*, *EMBO J.*, 2000, 19, 3159). One mechanism by which this can occur is over expression of the receptor at the protein level, for example as a result of gene amplification. This has been observed in many common human cancers (reviewed in Klapper *et al.*, *Adv. Cancer Res.*, 2000, 77, 25) such as, non-small cell lung cancers (NSCLCs) including adenocarcinomas (Cerny *et al.*, *Brit. J. Cancer*, 1986, 54, 265; Reubi *et al.*, *Int. J. Cancer*, 1990, 45, 269; Rusch *et al.*, *Cancer Research*, 1993, 53, 2379; Brabender *et al.*, *Clin. Cancer Res.*, 2001, 7, 1850) as well as other cancers of the lung (Hendler *et al.*, *Cancer Cells*, 1989, 7, 347).

It is now several decades since the study of retroviral mediated cellular transformation began to revolutionize our understanding of malignant transformation. Transformation was shown to be dependent on oncogenes carried by viruses and these were shown to have mammalian cellular counterparts, proto-oncogenes. In 1984, EGFR was described as the mammalian counterpart of the retroviral oncogene, v-erbB (Downward *et al.*). This, coupled to earlier observations describing a two component autocrine growth promoting mechanism in cancer cells consisting of EGF ligand and its receptor EGFR (Sporn & Todaro), strengthened

the hypothesis that EGFR signalling is an important contributor to tumourigenesis. Subsequent reports continued to provide evidence that EGFR is an attractive target for therapeutic intervention in Cancer (see Yarden & Sliwkowski for review). EGFR is markedly overexpressed across a large variety of epithelial Cancers (see Salomon et al) and some immunohistochemical studies have demonstrated EGFR expression is associated with poor prognosis. In addition to overexpression, it is recognised that there is potential for deregulated EGFR signalling in tumours via a number of alternative mechanisms including i) EGFR mutations ii) increased ligand expression and enhanced autocrine loop and iii) heterodimerisation and cross talk with other erbB receptor family members.

In addition, a wealth of pre-clinical information suggests that the erbB family of receptor tyrosine kinases are involved in cellular transformation. In addition to this, a number of pre-clinical studies have demonstrated that anti-proliferative effects can be induced by knocking out one or more erbB activities by small molecule inhibitors, dominant negatives or inhibitory antibodies (reviewed in Mendelsohn et al., Oncogene, 2000, 19, 6550).

Thus it has been recognised that inhibitors of these receptor tyrosine kinases should be of value as a selective inhibitor of mammalian cancer cells (Yaish et al. Science, 1988, 242, 933, Kolibaba et al, Biochimica et Biophysica Acta, 1997, 133, F217-F248; Al-Obeidi et al, 2000, Oncogene, 19, 5690-5701; Mendelsohn et al, 2000, Oncogene, 19, 6550-6565).

A number of small molecule inhibitors of erbB family of receptor tyrosine kinases are known, particularly inhibitors of EGF and erbB2 receptor tyrosine kinases. For example European Patent Application No. 0566226 and International Patent Applications WO 96/33980 and WO 97/30034 disclose that certain quinazoline derivatives which possess an anilino substituent at the 4-position possess EGFR tyrosine kinase inhibitory activity and are inhibitors of cancer tissue.

It has been disclosed by J R Woodburn et al. in Proc. Amer. Assoc. Cancer Research, 1997, 38, 633 and Pharmacol. Ther., 1999, 82, 241-250 that the compound N-(3-chloro-4-fluorophenyl)-7-methoxy-6-(3-morpholinopropoxy)quinazolin-4-amine is a potent EGFR tyrosine kinase inhibitor. This compound is also known as Iressa (registered trade mark), gefitinib (United States Adopted Name), by way of the code number ZD1839 and Chemical Abstracts Registry Number 184475-35-2. The compound is principally identified hereinafter as gefitinib.

Gefitinib was developed as an inhibitor of epidermal growth factor receptor-tyrosine kinase (EGFR-TK), which blocks signalling pathways responsible for driving proliferation, invasion, and survival of cancer cells (Wakeling, A.E., et al. *Cancer Res*, 2002, 62(20), p5749). Gefitinib has provided clinical validation of small molecule inhibitors of EGFR. Potent anti-tumour effects as well as rapid improvements in NSCLC-related symptoms and quality of life have been observed in clinical studies that enrolled patients with advanced NSCLC who did not respond to platinum-based chemotherapy. The Phase II 'IDEAL' trials demonstrated that single agent gefitinib resulted in objective anti-tumour activity, symptomatic improvement and limited toxicity in patients with advanced NSCLC and previously treated with cytotoxic chemotherapy (Fukuoka et al., Kris et al). Objective response rate (Complete Response + Partial Response) was 18.4% and 11.8% respectively in the IDEAL 1 and IDEAL 2 trials. The differences in response in these clinical trials has been attributed to different population groups in the two trials, predominantly Japanese in IDEAL 1 and a predominantly European-derived population in IDEAL 2. Beyond objective responses, additional patients experienced stable disease and / or symptom improvement meaning that approximately 50% of patients overall benefit from gefitinib. The tumour response data has been the basis of initial regulatory approvals of gefitinib in advanced NSCLC in several markets.

It is important to be able to understand the basis of response to anti-cancer therapeutic agents such as gefitinib since this would allow clinicians to maximise the benefit/risk ratio for each patient, potentially via the development of diagnostic tests to identify patients most likely to benefit from gefitinib treatment. An obvious candidate marker of response to gefitinib has been EGFR expression level. However, gefitinib inhibition of growth of some cancer-derived cell lines and tumour xenografts is not well correlated with the level of expression of EGFR. Furthermore, studies alongside the IDEAL trials demonstrated that EGFR protein expression as measured by IHC was not an accurate predictor of response to gefitinib (Bailey et al). Although there are now several additional hypotheses based on genetics, genomics, proteomics, biochemical and other studies, there is still no pre-treatment predictive biomarker of gefitinib response currently approved by regulatory authorities. Possibly the most significant recent breakthrough in understanding gefitinib response has come from recent data (Lynch et al, Paez et al) indicating that mutation in the EGFR kinase domain predicts gefitinib hypersensitivity in NSCLC patients. Hypersensitivity is a vague term but in this field is generally understood to mean patients experiencing objective tumour responses (i.e. marked tumour regression,

normally above 50%). As well as demonstrating the EGFR mechanism of action for gefitinib, this may provide a basis for venturing into other disease settings such as first line, adjuvant and possibly earlier cancer intervention with EGFR inhibitors in a targeted subpopulation in NSCLC patients and other types of cancers carrying the EGFR mutation.

However, it is likely that restricting prescription of gefitinib to the mutant EGFR carrying tumour subgroup will deprive many patients who could benefit from gefitinib. Firstly there are emerging reports of gefitinib hypersensitive patients with undetectable EGFR mutation in their tumour and other patients with EGFR mutation who do not respond to gefitinib. Secondly, data reported at ASCO 2004 (Shepherd et al) indicated that the EGFR small molecule tyrosine kinase inhibitor erlotinib (Roche, Genentech, OSI) prolongs survival in advanced NSCLC previously treated with chemotherapy, by ~2 months across the population with resulting 41% reduction in risk of death at one year. Most interestingly, the survival benefit appears to be derived from patients in the stable disease response population as well as hypersensitive patients. This highlights the likely importance of identifying likely gefitinib responsive patients beyond those carrying EGFR mutation. Definitive survival benefit is also likely to be demonstrated from ongoing clinical trials with gefitinib.

The differential response of patients to chemotherapy treatments indicates that there is a need to find methods of predicting which treatment regimes best suit a particular patient.

There is an increasing body of evidence that suggests that patients' responses to numerous drugs may be related to a patients' genetic, genomic, proteomic, biochemical or profile and that determination of the genetic factors that influence, for example, response to a particular drug could be used to provide a patient with a personalised treatment regime. Such personalised treatment regimes offer the potential to maximise therapeutic benefit to the patient, whilst minimising, for example side effects that may be associated with alternative and less effective treatment regimes.

Therefore there is a need for methods that can predict a patients' response to a drug based on the results of a test that indicates whether the patient is likely to respond to treatment or to be resistant to treatment.

The present invention is based on the discovery that the sensitivity of tumours to therapeutic agents can be predicted from the gene expression profile of the tumour and hence that the suitability of tumour patients for treatment with such therapeutic agents can be determined by measuring the relative expression levels of particular genes in tumour tissue.

According to one aspect of the present invention there is provided a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 as defined herein whereby to predict an increased likelihood of response to the erbB receptor drug.

According to another aspect of the present invention there is provided a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or DAPK2 whereby to predict an increased likelihood of response to the erbB receptor drug.

In one embodiment the method comprises testing a biological sample from the mammal for expression of any one of ACOX2, NPAS2, NES, CHST7, GSPT2, DAPK1, DAPK2 or TNNC1. More preferably the method comprises testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7 or DAPK1. More preferably the method comprises testing a biological sample from the mammal for expression of at least two of NPAS2, NES, CHST7 or DAPK1. More preferably the method comprises testing a biological sample from the mammal for expression of at least three of NPAS2, NES, CHST7 or DAPK1. More preferably still the method comprises testing a biological sample from the mammal for expression of NPAS2, NES, CHST7 and DAPK1.

In an alternative embodiment the method comprises testing a biological sample from the mammal for expression of any one of NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2, VAMP4, DAPK1, DAPK2, MLLT3, TNNC1 or KIAA0931. More preferably the method comprises testing a biological sample from the mammal for expression of any one of DAPK1, DAPK2 or NES. More preferably the method comprises testing a biological sample from the mammal for expression of at least two of DAPK1, DAPK2 or NES. More preferably the method comprises testing a biological sample from the mammal for expression of DAPK1, DAPK2 and NES.

In a preferred embodiment the method additionally comprises testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1, SLC20A1, SPRY2 or PGM1. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an alternative preferred embodiment the method additionally comprises testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1, HCA127, UBL5, ZNF23, UROD, CD44, SPRY1, RAPGEF2, SLC20A1, NRP1, PGM1, SPRY2, PTGER3, SCN10A, KITLG, CDH1, HOP, BCL3 or OLFM1. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

Preferably the tumour is selected from the group consisting of leukaemia, multiple myeloma, lymphoma, bile duct, bone, bladder, brain, CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head, neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural membrane, peritoneal membrane, prostate, renal, skin, testicular, thyroid, uterine and vulval. More preferably the tumour is selected from one of non-small cell lung, pancreatic, head or neck. More preferably the tumour is selected from one of non-small cell lung, head or neck.

Preferably the erbB receptor drug is selected from any one of gefitinib, erlotinib, PKI-166, EKB-569, HKI-272, lapatinib, canertinib, AEE788, XL647, BMS 5599626, cetuximab, matuzumab, panitumumab, MR1-1, IMC-11F8 or EGFR11. Most preferably the erbB receptor drug is gefitinib.

In a further preferred embodiment of the method of the invention the mammal is a human and the method comprises testing a biological sample from the human for increased expression of DAPK1 and decreased expression of NPAS2, NES, CHST7 or EMP1 whereby to predict an increased likelihood of response to gefitinib. In an alternative preferred embodiment of the method of the invention the mammal is a human and the method comprises testing a biological sample from the human for increased expression of DAPK1 and DAPK2 and decreased expression of NES and EMP1 whereby to predict an increased likelihood of response to gefitinib.

According to another aspect of the invention there is provided an isolated set of marker genes identified as having differential expression between tumour cells that are sensitive and resistant to an erbB receptor drug said gene set comprising one or more genes selected from at least the group consisting of the genes listed in Table 1 defined herein or DAPK2, including gene specific oligonucleotides derived from said genes. Preferably the set comprises at least 2

genes, more preferably at least 3 genes, more preferably at least 4 genes. More preferably the set comprises at least one gene selected from Table 2 as defined herein.

According to another aspect of the invention there is provided an isolated set of marker genes identified as having differential expression between tumour cells that are sensitive and resistant to an erbB receptor drug said gene set comprising one or more genes selected from at least the group consisting of the genes listed in Table 1 defined herein, including gene specific oligonucleotides derived from said genes. Preferably the set comprises at least 2 genes, more preferably at least 3 genes. More preferably the set comprises at least one gene selected from Table 2 as defined herein.

The present invention permits the improved selection of a patient, having or suspected of having a tumour, for treatment with an erbB receptor drug, in order to predict an increased likelihood of response to the erbB receptor drug.

In one embodiment, the method comprises testing a biological sample from the mammal for expression of at least one or more of the following from Table 1, which are found at lower levels in sensitive cells NPAS2, NES, CHST7, ACOX2 or GSPT2 or at least one or more of the following which are found at higher levels in sensitive cells DAPK1 or TNNC1. The Affymetrix ID and Affymetrix probe sequence for these genes are displayed in Table 1. In a preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of DAPK2 which is found at higher levels in sensitive cells, whereby to predict an increased likelihood of response to the erbB receptor drug.

In an alternative embodiment, the method comprises testing a biological sample from the mammal for expression of at least one or more of the following from Table 1, which are found at lower levels in sensitive cells NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2 or VAMP4 or at least one or more of the following which are found at higher levels in sensitive cells DAPK1, DAPK2, MLLT3, TNNC1 or KIAA0931. The Affymetrix ID and Affymetrix probe sequence for these genes are displayed in Table 1.

In a preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 2, whereby to predict an increased likelihood of response to the erbB receptor drug. In a preferred embodiment, the method comprises testing a biological sample from the mammal for expression of any one of the following genes listed in Table 2, which are found at lower levels in sensitive cells EMP1, SLC20A1, SPRY2 or PGM1, whereby to predict an increased likelihood of response to the

erbB receptor drug. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an alternative preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 2, whereby to predict an increased likelihood of response to the erbB receptor drug. In a preferred embodiment, the method comprises testing a biological sample from the mammal for expression of any one of the following genes listed in Table 2, which are found at lower levels in sensitive cells EMP1, HCA127, UBL5, ZNF23, UROD, CD44, SPRY1, RAPGEF2, SLC20A1, NRP1, PGM1 or SPRY2 or at least one or more of the following which are found at higher levels in sensitive cells PTGER3, SCN10A, KITLG, CDH1, HOP, BCL3 or OLFM1 whereby to predict an increased likelihood of response to the erbB receptor drug. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an especially preferred embodiment the method comprises testing a biological sample from the mammal for expression of NPAS2, NES, CHST7, DAPK1 and EMP1. High NPAS2, NES, CHST7 and EMP1 levels are associated with resistance to gefitinib and high DAPK1 levels are associated with sensitivity to gefitinib. Preferably, the assessment of expression comprises determination of whether DAPK1 levels are increased and NPAS2, NES, CHST7 and EMP1 levels are reduced.

In an alternative especially preferred embodiment the method comprises testing a biological sample from the mammal for expression of DAPK1, DAPK2, NES and EMP1. High EMP1 and NES levels are associated with resistance to gefitinib and high DAPK1 and DAPK2 levels are associated with sensitivity to gefitinib. Preferably, the assessment of expression comprises determination of whether DAPK1 and DAPK2 levels are increased and EMP1 and NES levels are reduced. In a most preferred embodiment the invention comprises determining the level of DAPK1 and EMP1.

According to another aspect of the invention there is provided a method for predicting clinical outcome of treatment with an erbB receptor drug for a mammal, having or suspected of having a tumour, comprising determining the level of any of the genes as described hereinabove in a biological sample taken from the tumour, or suspected tumour, wherein a poor outcome is predicted if:

- a) the expression level of DAPK1 is reduced; and /or

- b) the expression level of NPAS2, NES, CHST7 and EMP1 is increased.

According to another aspect of the invention there is provided a method for classifying cancer comprising, determining the level of any of the genes as described hereinabove in a biological sample taken from a tumour, or suspected tumour, wherein tumours expressing elevated levels of DAPK1 and / or reduced levels of NPAS2, NES, CHST7 or EMP1 are predicted as sensitive to treatment with erbB receptor drugs.

According to another aspect of the invention there is provided a method for predicting clinical outcome of treatment with an erbB receptor drug for a mammal, having or suspected of having a tumour, comprising determining the level of any of the genes as described hereinabove in a biological sample taken from the tumour, or suspected tumour, wherein a poor outcome is predicted if:

- a) the expression level of DAPK1 or DAPK2 is reduced; and /or
- b) the expression level of EMP1 or NES is increased.

According to another aspect of the invention there is provided a method for classifying cancer comprising, determining the level of any of the genes as described hereinabove in a biological sample taken from a tumour, or suspected tumour, wherein tumours expressing elevated levels of DAPK1 or DAPK2 and / or reduced levels of EMP1 or NES are predicted as sensitive to treatment with erbB receptor drugs.

According to another aspect of the invention there is provided a method for treating a disease condition in a mammal having, or suspected of having, a tumour, predicted to be resistant or non responsive to erbB receptor drug treatment based on the level of any of the genes as described hereinabove, comprising: providing a resistance-surmounting quantity of an erbB receptor drug and administering the resistance-surmounting quantity of the erbB receptor drug to the mammal.

In a preferred embodiment the mammal is a primate. In a most preferred embodiment the mammal is a human. In a preferred embodiment the patient is a primate. In a most preferred embodiment the patient is a human.

The term "erbB receptor drug" includes drugs acting upon the erbB family of receptor tyrosine kinases, which include EGFR, erbB2 (HER), erbB3 and erbB4 as described in the background to the invention above. In a preferred embodiment the erbB receptor drug is an erbB receptor tyrosine kinase inhibitor. In a preferred embodiment the erbB receptor drug is an EGFR tyrosine kinase inhibitor.

In a more preferred embodiment the EGF receptor tyrosine kinase inhibitor is selected from gefitinib, Erlotinib (OSI-774, CP-358774), PKI-166, EKB-569, HKI-272 (WAY-177820), lapatinib (GW2016, GW-572016), canertinib (CI-1033, PD183805), AEE788, XL647, BMS 5599626 or any of the compounds as disclosed in WO03/082831, WO05/012290, WO05/026157, WO05/026150, WO05/026156, WO05/028470, WO05/028469, WO2004/006846, WO03082831, WO03/082290 or PCT/GB2005/000237.

In another preferred embodiment the erbB receptor drug is an anti-EGFR antibody such as for example one of cetuximab (C225), matuzumab (EMD-72000), panitumumab (ABX-EGF/rHuMAb-EGFr), MR1-1, IMC-11F8 or EGFR11.

We contemplate that erbB receptor drugs may be used as monotherapy or in combination with other drugs of the same or different classes. In an especially preferred embodiment the EGF receptor tyrosine kinase inhibitor is gefitinib.

In a preferred embodiment the present invention is particularly suitable for use in predicting the response to the erbB receptor drug as described hereinbefore in those patients or patient population with a tumour which is dependent alone, or in part, on an erbB tyrosine kinase receptor. Such tumours include, for example, non-solid tumours such as leukaemia, multiple myeloma or lymphoma, and also solid tumours, for example bile duct, bone, bladder, brain/CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head and neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural/peritoneal membranes, prostate, renal, skin, testicular, thyroid, uterine and vulval tumours.

In a preferred embodiment the present invention is particularly suitable for identifying a patient with head, neck, pancreatic, glioblastoma, colorectal or breast tumour for drug treatment. In an especially preferred embodiment the present invention also is particularly suitable for identifying those patients with NSCLC, more particularly advanced NSCLC including advanced adenocarcinoma that will respond to treatment with an erbB receptor drug as hereinbefore defined.

The present invention provides advantage in the treatment of tumours such as NSCLC, especially advanced NSCLC by identifying "individual cancer profiles" of NSCLC and so determining which tumours would respond to erbB receptor drug such as gefitinib.

The present invention is particularly useful in the treatment of patients with advanced NSCLC who have failed previous chemotherapy, such as platinum-based chemotherapy. The present invention is also particularly useful in the treatment of patients with locally advanced

(stage IIIB) or metastasized (stage IV) NSCLC who have received previous chemotherapy, such as platinum-based chemotherapy. The present invention is also useful in adjuvant therapy or as a first-line therapy.

In a preferred embodiment there is provided a method of selecting a human, having or suspected of having a tumour, for treatment with gefitinib which comprises testing a biological sample, from the mammal for expression of NPAS2, NES, CHST7, DAPK1 and EMP1, whereby to predict an increased likelihood of response to gefitinib.

In a preferred embodiment there is provided a method of selecting a human, having or suspected of having a tumour, for treatment with gefitinib which comprises testing a biological sample, from the mammal for expression of DAPK1, DAPK2, NES and EMP1 whereby to predict an increased likelihood of response to gefitinib.

According to another aspect of the invention there is provided a method of predicting the responsiveness of a patient or patient population with cancer, for example lung cancer, to treatment with chemotherapeutic agents, especially erbB receptor drugs, comprising comparing the differential expression of any of the genes described herein.

In one embodiment the assessment of expression is performed by gene expression profiling using oligonucleotide-based arrays or cDNA-based arrays of any type, particularly where large numbers of genes are analysed simultaneously. In an alternative embodiment, RT-PCR (reverse transcription- Polymerase Chain Reaction), real-time PCR, *in-situ* hybridisation, Northern blotting, Serial analysis of gene expression (SAGE) for example as described by Velculescu et al Science 270 (5235): 484-487, or differential display or any other method of measuring gene expression at the RNA level could be used. Details of these and other general molecular biology techniques can be found in Current Protocols in Molecular Biology Volumes 1-3, edited by F M Ausubel, R Brent and R E Kingston; published by John Wiley, 1998 and Sambrook, J. and Russell, D.W., Molecular Cloning: A Laboratory Manual, the third edition, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York, 2001.

In another embodiment the assessment of expression is performed by measurement of protein levels encoded by the aforementioned genes. For example, an immunohistochemistry-based assay or application of an alternative proteomics methodology.

In another embodiment the assessment of expression is performed by measurement of activity of the proteins encoded by the aforementioned genes, for example in a bioassay.

In a preferred embodiment the biological sample would have been obtained using a

minimally invasive technique to obtain a small sample of tumour, or suspected tumour, from which to determine gene expression profile. Such techniques include, for example tumour biopsy, such as transbronchial biopsy. The profile of gene expression of transbronchial biopsy specimens whose size is about 1 mm may be measured for example using a suitable amplification procedure.

Another aspect of the invention provides a kit for use in a method of predicting the responsiveness of a patient or patient population with a tumour, to treatment with chemotherapeutic agents, especially erbB receptor drugs, comprising a means for measuring the levels of any of the genes as described hereinabove. Preferably the genes are attached to a support material or membrane such as nitrocellulose, or nylon or a plastic film or slide.

In a further preferred embodiment the present invention includes administration of an erbB receptor drug to a mammal selected according the methods described hereinabove.

According to another aspect of the invention there is provided a method of using the results of the methods described above in determining an appropriate dosage of an erbB receptor drug.

In a preferred embodiment the biological sample comprises either a single sample which may be tested for expression of any of the genes as described hereinabove, or multiple samples which may be tested for expression of one or more of the genes as described hereinabove.

The invention is illustrated by the following non-limiting examples in which:

Fig 1 illustrates a xenograft (A549 cell line) which when grown as a xenograft in athymic mice is sensitive to gefitinib. This involved oral dosing, once daily, at the dose indicated. Y axis = mean tumour volume in cm^3 ; x axis = days after treatment.

Fig 2 illustrates a xenograft (MKN45 cell line) which when grown as a xenograft in athymic mice is resistant to gefitinib. This involved oral dosing, once daily, at the dose indicated. Y axis = mean tumour volume in cm^3 ; x axis = days after treatment.

Figures 3, 4, 5 and 6 show examples of specific gene expression profiled across a wider panel of gefitinib sensitive and resistant lines, where definition of sensitivity is based on response to gefitinib when grown as a xenograft, to increase confidence that the expression profile of each gene is truly predictive. Iressa sensitivity is based on xenografts data. The cell lines and the tumours from which they are derived are as follows; KB – head and neck, HT29 - colon, BT474 – breast, DU145 – prostate, LoVo – colon, MCF7 – breast, GEO – colon, A549 – lung,

A431 - epidermoid, H322 - lung, HX147 - lung, RT112 - bladder, MiaPaCa2 - pancreas, MKN45 - gastric, MDAMB231 - breast, PC3 - prostate, Calu6 - lung, SW620 - colon.

The legend key is S=sensitive, U=unknown and R=resistant.

Fig 3 shows EMP1 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 4 shows DAPK1 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 5 shows DAPK2 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 6 shows NES basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Example 1

Gene Expression in Gefitinib Resistant or Sensitive Tumour Cell Lines – Cell Culture and Xenograft Studies

We identified genes useful to predict response to erbB receptor drugs in the clinic. This is based on studies with gefitinib, but the findings are applicable to erbB receptor drugs in general.

The gene lists have been assembled by comparing tumour cell lines which have been demonstrated to be either sensitive to gefitinib or resistant to gefitinib. This definition is based on the response observed when the tumour cell line is implanted into nude mice and grown as a xenograft. This definition has been used for all the pre-clinical studies described herein.

Initially a small panel of six human tumour cell lines were assembled, three which are sensitive to gefitinib and three which are resistant to gefitinib in the xenograft setting defined above.

The sensitive cell lines were;

1. Lovo (ATCC¹ No. CCL-229) – colon tumour cell line
2. KB (ATCC No. CCL-17) – initially reported as a nasopharyngeal cell line (although more recently reported as Hela derived (cervical carcinoma))
3. HT29 (ATCC No. HTB-38) – colon tumour cell line

The resistant cell lines were;

1. MKN 45 (source - Nottingham University, UK) – gastric tumour cell line
2. Calu 6 (ATCC No. HTB-56) – lung tumour cell line
3. PC3 (ATCC No. CRL-1435) – prostate tumour cell line

¹ATCC = American Type Culture Collection

The cell lines were grown both in cell culture and as xenografts, RNA prepared and the basal expression profiles determined by measuring RNA expression on the Affymetrix microarray platform. As part of our studies, the term 'basal' has been used to indicate constitutive or steady state expression levels (rather than expression levels which are modulated as a consequence of administration of an erbB ligand or gefitinib to the cells). Figure 1 illustrates the sensitivity of A549 xenografts (used in Example 3 below) to treatment with gefitinib. Figure 2 illustrates the resistance of MKN45 xenografts to gefitinib. See Example 2 below for analysis of results.

Example 2

Statistical analyses of cell culture and xenograft data sets

The following statistical analyses were performed separately for cell culture and xenograft data sets. Probe sets were eliminated if their signal was not distinguishable from background noise across all RNA samples in the set. Mixed ANOVA (see for example Scheffe, 1959) was applied separately to each remaining probe set to generate p values. The p values were then used to calculate Q values (Storey). The Q values indicate the expected proportion of genes in a gene list which are not truly differentially expressed but have been falsely discovered (False Discovery Rate or FDR). Q value cut-offs appropriate in the different studies were identified and applied, based on graphical examination of the p value and Q value results, in conjunction with fold change. The final genelists for each study were generated using Q value and fold change (FC) cut-offs. The different genelists were then combined to display an overall list of genes which showed consistent differences in expression profiles between the cell lines in the sensitive and resistant groups.

Further details of the analysis procedures are provided as follows. Fold change (FC) was calculated based on the mean of sensitive cells divided by the mean of resistant cells. To generate gene lists, FC cut-off of two-fold (2X) change in either direction was used in all cases. Furthermore FDR Q values were used to narrow down the lists and obtain the most significant gene changes across sensitive versus resistant cell lines. In the case of cell culture, Q value cut-off is 0.3. In the case of xenograft, Q value cut-off is 0.6. The different cut-offs used reflect the different design and variance values associated with each experiment.

In cell culture studies, lists were obtained based on the above criteria for cells grown either in full serum containing medium or in charcoal stripped serum. In the xenograft study, the same as above was performed for separate sets of tumours harvested at 18hr intervals. Gene lists contain some redundancy in genes where appropriate to illustrate consistency of results obtained for example with different probe sets.

Example 3

Identification of predictive genes

Genes which have not previously been identified as predictive of erbB receptor drug sensitivity are listed in Table 1. Other genes which we have identified to be optionally used in combination with Table 1 genes are listed in Table 2.

Key to Tables:

'Affymetrix ID' – the Affymetrix probe set identifier

'Sequence' – target sequence relating to the Affymetrix probe set indicated by 'Affymetrix ID'

"+" if up in sensitive" means that the gene is relatively highly expressed in sensitive cells. (Consequently, absence of a "+" means that the gene is relatively highly expressed in resistant cells).

'Gene Title' - The current annotation of the gene relating to 'Affymetrix ID' based on UniGene
133

'Gene Symbol' – shorthand synonym for the gene title

'Locus Link' & RefSeq Transcript ID' are provided for gene identification purposes.

Combining genes has the potential to generate an improved diagnostic over genes used in isolation. Collective gene expression profiles (at the RNA and/ or protein level) may be more likely to identify patients most likely to benefit from gefitinib rather than the expression level of one gene in isolation.

It may be more practical when developing a pre-treatment response prediction diagnostic to work with a truncated gene list from tables 1 and / or 2. A number of criteria have been used to shorten the gene list to identify those genes which are most predictive of response. Firstly the statistical (p values and Q values or FDR values) can indicate the statistical significance of a gene.

Secondly, the differential expression (fold change) between the sensitive and resistant groups indicates the potential sensitivity of a marker to be used in a diagnostic test (highest fold change between sensitive group and resistant group is preferred).

Thirdly, we have performed RT-PCR based expression profiling across a wider panel of gefitinib sensitive and resistant human tumour cell lines to increase confidence that the expression profile of each gene is truly predictive. Figs 3, 4, 5 and 6 show examples of specific gene expression profiled across a wider panel of cell lines as set out below.

The sensitive human tumour cell lines, where definition of sensitivity is based on response to Iressa when grown as a xenograft:

- a. BT474 (ATCC No. HTB-20) – breast tumour cell line
- b. DU145 (ATCC No. HTB-81) – colon tumour cell line

- c. MCF7 (ATCC No. HTB-22, sourced from ICRF (now CR-UK), London), - breast tumour cell line
- d. GEO colon tumour cell line. RNA obtained from Fortunato Ciardiello, Cattedra di Oncologia Medica, Dipartimento Medico-Chirurgico di Internistica Clinica e Sperimentale "F. Magrassi e A. Lanzara, " Seconda Universita delgi Studi di Napoli, Via S. Pansini, 5-80131, Naples, Italy.
- e. A549 (ATCC No. CCL-185) – lung tumour cell line
- f. A431 (ATCC No. CRL-155) – epidermoid cell line

The resistant human tumour cell lines, where definition of sensitivity is based on response to Iressa when grown as a xenograft:

- 1) HX147 - (source: ICRF (now CR-UK), London) – lung tumour cell line
- 2) RT112 - bladder tumour cell line (DSMZ No ACC 418)
- 3) MiaPac2 (ECACC 85062806, ref. no. 001611) pancreatic tumour cell line
- 4) MDAMB231 (ATCC No. HTB-26) – breast tumour cell line
- 5) SW620 (ECACC CCL-227) – colon tumour cell line

ATCC = American Type Culture Collection

DSMZ - Deutsche Sammlung von Mikroorganismen und Zellkulturen GmbH (German Collection of Micro-organisms and Cell Cultures)

ECACC = European Collection of Cell Cultures

In isolation, each of these genes is reasonably predictive of gefitinib response, but collectively they can be applied to make predictions with a higher level of confidence.

The Affymetrix probe sets identifiers for the genes in the above diagnostic genelists are indicated in Tables 1 and 2. Current Affy IDs are based on Affy U133 chipset. For the avoidance of doubt, the target sequences of the Affymetrix probe sets which identified the listed genes are also provided in Tables 1 and 2.

Without wishing to be bound by theoretical considerations, it is contemplated that the specific sequences used to detect target genes in the Examples may define specific splice variants or sequences in homologous genes. Therefore in one embodiment, a listed gene for use in the method of the invention is defined by the specific sequence used in said Examples. In another embodiment, a gene for use in the method of the invention is not limited by the specific sequence used in these Examples. Indeed the fact that some genes in Tables 1 and 2 have been identified using different sequences (gene "redundancy") and confirmatory RT-PCR studies (see

Example 4) provides evidence that usefulness in the method of the invention is not generally limited to the specific sequences used to measure the target gene.

Note, in the event of a discrepancy in the sequence between Tables 1 and 2 and the Sequence Listing, the sequence as provided in the Tables is preferred.

Table 1: as described in priority application US60/619027 filed on 18/10/2004.

Gene Symbol	Gene Title	Affymatrix ID	+ up in sensitive	Sequence	LocusLink	RefSeq Transcript ID	SEQ ID NO.
ACOX2	"acyl-Coenzyme A oxidase 2, branched chain / acyl-Coenzyme A oxidase 2, branched chain"	205364_at		Gtgcagcaattitacagaccctgaccccaatccggagctgaccacgacgagggcttggaa accagcaactgtctatcaactccaggctgtcaagggtgcaactgtctactatgtcacgtg aagggtttacagaaagctctggagaaactgaaaaatgaaccagcgattcagcaggt gtcacaagcgctctgtgacctccatgccatacatgaaatctgtgaciacactcgggtgact ttctccatgacgcctccctctgtctgttgcacaagtggacatggcaagaacagcctacgtg gacctgtccgcctgatccggaaagatgccatccctgtaaacgtatgtttgtgacttcacc gatcagtttaaatcaagccctggctgtatgatggaacgctctacgaacgctgttcc agtgggctcagaagctc	8309	NM_003500	SEQ ID NO:1
ACTR2	ARP2 actin-related protein 2 homolog (Yeast)	200729_s_at		gagctaaagctcgtgtttgttaatgctctgtttatccagaagcattaaaggtiaaccocat tgcacaagctatcaattctgcaaatattctttataataactgaccagtgcttaataaacaag caggctactcaaaataactgtgcagagtggtggttaataatgtgtgttaaaaaataacatlg gaatacaggaactgttgccaattgggtlaatttcatlgtgtttgtttgtattgtgaac ctggaaatacagaaaaattgactgttaaaaatgttggccaaaaataatcagaatttaatt tttttaattgtactgaaaacctatcaataactgttaattctcagccaactcttgaagctgaaa gaagactctgtgtattttglaaacggttagcagacacttctgccaagtgtcagaaaaatccaa tttatgaatcctctgcggtaactctgtgtatcgaaaaaataaccaaaatgtaaccatacaltg agttattctaa	10097	NM_005722	SEQ ID NO:2
APOL1	"apolipoprotein L1 / apolipoprotein L1"	209546_s_at	+	agaatagagaggaggtgaaggaacacagcaalgagaaggccaggaagaagaaagaa agagctgaaatggagaaagcccaagaggttagaacaggttgatacaggagaaga aaacagcggtccactacagaccacgcccaggttcaatgtctcccgaaagaatgaag tcttctcctgtgtgagtgccctgcccctgtcttcacagatccactcctcctgtctccgtgg ggccatctatcagtcagcagcggtctcctgaltgctcgttgggtgtgtgaatgtg atgggtccctccaggttactaaaagggtgcaatgtccctgcttgaacactgaaggcgag gtgtgt	8542	NM_003661 / NM_145343 / NM_145344	SEQ ID NO:3
C10orf10	chromosome 10 open reading frame 10 / chromosome 10 open reading frame 10	209183_s_at		aactcatcgtctgtgtgcaattgggagagttcccccattgatgaggggccaagata gaatctgtaccactcagctaccatcccccactacacccactccacacaggggc ctcatgtccatgtcaggggtcccccgtgtatgttgagagcagggccactgtccagctgtgc cacgtgggaagctcaagatgtcttaaggcccaaggcagggcacttggagcttgaagg acctatgtccatgagccatctgccaagaagaaggtgagggcatcaggagctcgaacgga atcagggctgggaadgacagaggtgaaagggacagagagagagagagagaggaaga ttgagctggggccaacagccaagctaccctggcaggtctctgccactcctgtct gtgagctgcaagctaggttaattcttttttgggtatttttaattgtcttgattgttaaatg ttttctgtctctttaaagctgttt	11067	NM_007021	SEQ ID NO:4

CALM1	"calmodulin 1 (phosphorylase kinase, delta)"	211984_at		tgacggctgicaglagccacgcgcgctgctgctgcgcagaggagaaatagcttaca cgaacccacagatcaggaaatctctcgaacaacgaatgaaattggccaagt ctactctcgctcaatcaattatgaacattgaattatgatactcaaatattct ccttgagcagatctccacaaatgctctaattctatctccacagcggcagttac tgcttttaattcccccgaagctgctcglaggagacagaaaattcttgctgctgataccctt ggaglaa	801	NM_006888	SEQ ID NO:5
CALM1	"calmodulin 1 (phosphorylase kinase, delta)"	211985_s_at		gagcgaatgagatcgaatattcagatgggcctttgagcactgttcgcaagggaaggc ttttctgattttgacaaatgaattttgcaacactcattggctcttcggcaacttacaca cattgaaat	801	NM_006888	SEQ ID NO:6
CD44	CD44 antigen (homing function and Indian blood group system)	210916_s_at		caagtgttgaggacgcagcctggggactctgcccctgcccctgagccctggcgca gactgattgataataacctgcgccttgaggctgattccacgcgggagaaaaatggctc gtacacgactctcgcagcgaggccgcctgacccctgcaaggccttcaatagcaactt ggccacatggccacagatggagaagctctgacatcgattgagacactgagctt ctctgcagtcacacagctgaagaaaggctggggcagagaagaaaaagctagtgatcaa cagtgcaatggagctgctggaggaacgaagcccaagtggaactcaacggagagc cagcaagctcagggaaaggctgattggtagaacaaggagctcgaagaactccag accagt	960	NM_000610	SEQ ID NO:7
CD44	CD44 antigen (homing function and Indian blood group system)	212063_at		attgaaacttttgctctcctgaagactccctaaataatgctcctgagtgaaaaatcaa aagagacaaaagacatctcgaatccattcaagcctgaglaaatggctttctag gaacactttccaaaagtttattgagattcaatacaacaccaaagaattgctttgag cacaactctcaactctgtatcacagaggagtaggagagaggaacacattgactta tctggaaaagcaaaatgtaacttaagaalaagaataacatggctcaactcattgta tagatatctgttgaaactcattgtttgagtttcaaaagaalagcccattgtcattctgt gcgtacaatgaccactgtattgtacttgcacttttcagagcacaccc	960	NM_000610	SEQ ID NO:8
CDS2	CDP- diacylglycerol synthase (phosphatidate cytidyltransferase) 2	212864_at		ttctatgcatccaccacaaaatccgcagaatgtaagtaagctcgtttataagatgg gttbaacctctcgcagacgtgaaagtttcagttttatttttttcagaaaagcagcaaaat tatttataatgctcggagaaaacacacactgtaatttcaagatgtaagcagtagaatg tactgtaactgagcccttccacatctgactgagcccaatgctcctcctgaggtccactaa ctgctgttttcaggacaaatgcacbaatgtttgtgcgtagacactgctgctgtaact ttctggggactcttcacatcgggcagggagagagggctctctgttcaatgcaaccttggc tgaecacccactgtagctgctgtgtgtgtatataattcttaagaggaggtgtgtgtgt gtgttttaaaagtcactattcttccagtgatttcaattgcacccatgactctcaataa accacaaagtctcgtctaaacctatggaaaacctaacctgatttagaccttgac	8760	NM_003818	SEQ ID NO:9

CHST7	carbohydrate (N-acetylglucosamine 6-O) / sulfotransferase 7 (N-acetylglucosamine 6-O) sulfotransferase 7	206756_at		56548	NM_019886	SEQ ID NO: 10
COMM D3	COMM domain containing 3	218048_at		23412	NM_012071	SEQ ID NO: 11
CUL2	cullin 2 / cullin 2	203078_at		8453	NM_003591	SEQ ID NO: 12
DAPK1	death-associated protein kinase 1 / death-associated protein kinase 1	203139_at	+	1612	NM_004938	SEQ ID NO: 13
DNAJC3	"DnaJ (Hsp40) homolog, subfamily C, member 3"	208499_s_at		5611	NM_006260	SEQ ID NO: 14

DPYSL 3	dihydropyrimidina se-like 3 / dihydropyrimidina se-like 3	201431_s _at		1809	NM_001387	SEQ ID NO:15
DUSP4	dual specificity phosphatase 4 / dual specificity phosphatase 4	204015_s _at		1846	NM_001394 / NM_057158	SEQ ID NO:16
EIF3S4	"eukaryotic translation initiation factor 3, subunit 4 delta, 44kDa / eukaryotic translation initiation factor 3, subunit 4 delta, 44kDa"	208887_at		8666	NM_003755	SEQ ID NO:17
EIF5A	eukaryotic translation initiation factor 5A	213757_at		1984	NM_001970	SEQ ID NO:18
FADS2	fatty acid desaturase 2 / fatty acid desaturase 2	202218_s _at		9415	NM_004265	SEQ ID NO:19

FLJ12442	hypothetical protein FLJ12442	218051_s_at			64943	NM_022908	SEQ ID NO:20
FLJ22028	hypothetical protein FLJ22028	213878_at	+		79912	NM_024854	SEQ ID NO:21
FNTB	"James/transferase, CAAX box, beta"	204764_at			2342	NM_002028	SEQ ID NO:22
GPCR5B	"G protein-coupled receptor, family C, group 5, member B / G protein-coupled receptor, family C, group 5, member B"	203632_s_at			51704	NM_016235	SEQ ID NO:23

[illegible]

KIAA0931	KIAA0931 protein	213407_at	+	attatgctccaaagccattcagtgatgcttcagacacacatataaggactgctagtgctcadt tttactctctcgggagggtccgactcccaatcatgaaggcaagtaacitccca gttagtgcatttgcoccatagtggggaanccactcctagattgagaaaaagagct acagcaatccctgctgttgcctcatttggatcagtcagtcacacataaagctctgt attcaaatctatgcacttccagatgctatagggtttctctcactgttgccaatggat gtcaaccagacagtggtgcatactacaggtttgtgc	23035	---	SEQ ID NO:29
KLHL7	kelch-like 7 (Drosophila)	220239_at		agtgtacagagcctccagagtggtgatgctttccactgtgtgatccctagtgga catgaatgaacgctccagatgttggcagtgagccacccctatctgcaggaatacgtcc aagacccacagtgaaigccggaacgacagatagctgaalccataataatactgtgt tttatgatacatcatgcctatgatgaagt	55975	NM_018846	SEQ ID NO:30
LAMC2	"laminin, gamma 2 / laminin, gamma 2"	202267_at		aagaaatgttctactcacactcagctgggtacacatccatccctccatccatcc atccatcttccatccattaccctccatccatccctccacacataattattgaglacctcgt gtgccaggggtgtgggacagtggtgacatagctcctccatcagagtgattgtgc tagtgaggaagacacagcatcttttaaaaaataaattaaacttacaadttgtgtcac aagtggtttattgcaataaccgtgtgtttgcaacctcttgcacacagacataatgtt gcaagacccctccatggggacactgtgtttggcaaggtgcacagagctctgggttg tgcacattcttgcattccacagctgtcacctgtgccttctacacactgatgcaacagact gtgtgattgatcaacacagtggtgaatgtcgtggaggaacacagaggtccactccact ggcgggaagacatgtgtgtgccttgc	3918	NM_005562 / NM_018891	SEQ ID NO:31
MLLT3	"myeloid/lymphoi d or mixed- lineage leukemia (trithorax homolog, Drosophila); translocated to, 3 / myeloid/lymphoid or mixed-lineage leukemia (trithorax homolog, Drosophila); translocated to, 3"	204918_s _at	+	aaggcatccacaggaatcatttaaaaaaagaatctgtcctgtttctaaaaa aaaaaacctgttgagaaattcttaatttggatctattattagtcagagtttcagcttctc agctgccagtggttactcatcttctccaaatctggaatcagagattttgtgttca catagtattctctagacacatttataatgaaaaaataaaatcttcttggggaacaaat ctgtgtattctgccataacagattatgtatataactgtatgacagtggttcaataactgtt agtgtctgtataatctccagaggtattctgttattgttggaagacgtgtgggtggg gggtattttgtctgtgttgcacctgtttgaacacagaaatctcctgtgtggaatgcaaa agaaagcaaatatttttaaaagaaaaaaacaaagtaactttgtgtcattatccatc ttctcca	4300	NM_004529	SEQ ID NO:32
MNAT1	menage a trois 1 (CAK assembly factor) / menage a trois 1 (CAK assembly factor)	203565_s _at		ccaagccactgcagatagacataggaacacatgttctcgtgactgtgagatgctagg aagactgtgtatttaaacatgacagctgcctccaccacagacactgtctggagggc tatactctctctgtgtgcacagacacacagatgcattcagtggtgtttctgtgca ggccagttaaccaattalaagattggacctgtggagctgaacacagggagagcaaa agtataagcagactataaaattatagctatgtcagctgcacacacagctcttccact agcagctgtgttaa	4331	NM_002431	SEQ ID NO:33

[illegible]

PCDH GC3	"protocadherin gamma subfamily C, 3"	209079_x _at	caagaagctcagcccaagatggggcttctcaacagggccctgcctctcgaagc ctcagctctcaccctgcaggagcgcttctctccgtgaaggccactgccaggccoc cagtgccgcccctagtgccatagcctggttaagttccaggtccctctctgcaata gacctctctccacccctctgcctcgggctcccgccatccaggggctgcca gagaacccagacccctgcctacagtagtgagcccccctctctctcggctgggt agaatagccagtagtgagtggggtgcttttacgtagtggggtgggca-gcgggc ggcggtccgcgagccgtctgctcgaicggccggcgcccggtgtgtgtttg tgctgtccacggcgttaagggaacccctcccgctac(gactctctctataagcgct ctctgcacatagtcacgtacgtccaccccccctctctctgctcacgcaagttaa ggatgggctctctcaacaggggccctgcctctgaagcctcagctcctcactggc agggtccgttctctccgtgaaggccactgccaggtcccgagtgcccccctagtg gccatagcctggttaagttccagtgccctctgtgcatagaacctctctccacccc ctctgcccctgggtcccgccatccagggggtgcagagaacccccaagcttg ccctacagtagtgtagggcccccctctctcggctgtgtagaataagcaatagtg agtggtgtgttttacgtagtggggtggcgagggcgccggcgctccggca ggcgtctctgactgacggcgggcgccgtgtgtgtgtgtgtgtgtgtgtgtgt aaggcgaacccctcccgctac(gactctctataagcgctctctctcgaatagcaag tagctccaccccccctctctctgctcagcgaagtgtttatactctatattatggc tttttctcgacaa	5098	NM_002588 / NM_032402 / NM_032403	SEQ ID NO:38
PCDH GC3	"protocadherin gamma subfamily C, 3 / protocadherin gamma subfamily C, 3"	211066_x _at	ggcagcttgggtgagctaacaggaccaatggattaaactggcatttaagtcacag gaagctcgaagcaggttaggaacagggtccctctgagaggtcagaggggctctgt gggtgctgggtacccaggtgcacgtggaagggtcagcgagggccagtgcc ctcctgtgcatagaacctctctccacccctctgcccctgggtcccgccatccag cggggtgccagagaacccccaagacctgacctacagtagtgagtggtgtgtgtgt tttcgggtggtgagaatagccagtagtgtagtggtgtgtgtgtgtgtgtgtgtgt ggcagcgggcgggcggtcccgccgagccgtgctcctgactgcccggcgggc ccgt tccataagcgctctctctgcatagtcagctcagctcagctccaccccccctctctgctc acgcaagttaa	5098	NM_002588 / NM_032402 / NM_032403	SEQ ID NO:39
PCDH GC3	"protocadherin gamma subfamily C, 3 / protocadherin gamma subfamily C, 3"	215836_s _at	ggcagcttgggtgagctaacaggaccaatggattaaactggcatttaagtcacag gaagctcgaagcaggttaggaacagggtccctctgagaggtcagaggggctctgt gggtgctgggtacccaggtgcacgtggaagggtcagcgagggccagtgcc ctcctgtgcatagaacctctctccacccctctgcccctgggtcccgccatccag cggggtgccagagaacccccaagacctgacctacagtagtgagtggtgtgtgtgt tttcgggtggtgagaatagccagtagtgtagtggtgtgtgtgtgtgtgtgtgtgt ggcagcgggcgggcggtcccgccgagccgtgctcctgactgcccggcgggc ccgt tccataagcgctctctctgcatagtcagctcagctcagctccaccccccctctctgctc acgcaagttaa	5098	NM_002588 / NM_032402 / NM_032403	SEQ ID NO:40
PDLM 7	PDZ and LIM domain 7 (enigma) / PDZ and LIM domain 7 (enigma)	203370_s _at	ggcagcttgggtgagctaacaggaccaatggattaaactggcatttaagtcacag gaagctcgaagcaggttaggaacagggtccctctgagaggtcagaggggctctgt gggtgctgggtacccaggtgcacgtggaagggtcagcgagggccagtgcc ctcctgtgcatagaacctctctccacccctctgcccctgggtcccgccatccag cggggtgccagagaacccccaagacctgacctacagtagtgagtggtgtgtgtgt tttcgggtggtgagaatagccagtagtgtagtggtgtgtgtgtgtgtgtgtgtgt ggcagcgggcgggcggtcccgccgagccgtgctcctgactgcccggcgggc ccgt tccataagcgctctctctgcatagtcagctcagctcagctccaccccccctctctgctc acgcaagttaa	9260	NM_005451 / NM_203352 / NM_203353 / NM_213636	SEQ ID NO:41

PEX3	peroxisomal biogenesis factor 3	203972_s _at	tggatccaaacatttattgcaattatattgattgcaagatgaagaaactccattagcagtg gcaggccgtggacgttctctcagacattaccattaaacttccatgaacttag agacattgtgaaagccagatttagtgcacattgaatactgtttaaacccaggtttt agtagactctagacaattggctggtcttctgacactgaacagaccctgcaaca tggtaactctatgaattgcttccagtcagccctgcttttagtaagataattccaala giaacggacagalcacttccagttgcagtaaacacctagtcatttggtaaggatctg ttgacaattggagcaagtgaaagacttgcgtcaatggtatgaagcttttagtacct cagcaactggagaaat	8504	NM_003630	SEQ ID NO: 42
PIN1	protein (peptidyl- prolyl cis/trans isomerase) NIMA-interacting 1 / protein (peptidyl-prolyl cis/trans isomerase) NIMA-interacting 1	202927_at	agccattgaagacgcctctgttgcgtgcggacgggggagatgagcgggcccgtgt tcacggattccggcatccacatcctccgcactgagtgagggtggggagccag gccctggctcggggcagggcagggggtagggcggccagctcccttgcctgc cagccagtgccgaaccccccactccctgccaccgtcacacagattattgtccca caatggctggggggggccctccagattggggccctggggcccccactccctgic calcccagttggggtgcgacccgacgattcccttaaggaattgactcagcagg gggggaggtccacagaccagggcagtggtgggggggtgttcccaagagaa ggcctgtcagcagagcccccgtgtccccccaggtgtggagggcagacacag ggccgaattgttctagtgaggccacgctcctgttcagtcgcaagggaacactcat gcggcagcc	5300	NM_006221	SEQ ID NO: 43
PRKCA	"protein kinase C, alpha"	213093_at	gattaaacgactgtgttgcacctctgttaactttaggagatccactcctgtgattgt agaonttggatattctctggaggaataatcattctttcttgaagggttggttactaga aattcaaatcaatcaagaaggcagttactatttgaatcaagggtttctaaatfa acctcacatccctctgttgggtcttcaagaatactttataaacagagcatttgaagt cattgtttgtctaatgattgtgtgtgagggacataccacgttttaaatcatttaattgaa aaacatataagcccaactttgttggggagagagacgggaggttgggttttctt ctgtataagccactactgacaaaatgtagggccattcaaccgcaaacaccatttgg ttatatcgacagaggagacggatgtgaaattactgcatgtcttttttttcagtttgataacc tcaatctccgttgcagatcacgtttgttagaa	5578	NM_002737	SEQ ID NO: 44
RIOK3	RIO Kinase 3 (yeast)	202129_s _at	tgaatgacgttggccatgctgaacctcagtgagataacatgctgtggcatgctggaa aggctgtgtgactcagtcagtcagtagaacctaccacccctcagccctggga gttctgtccgggactgcaggaaatgtctcgcagttttccagaaaggaggagcagg aagccctagtgaacgagaactctcaatgctgtttcaggcttaaacatcacagcagat aatgaagctgatttttagctgagatagaagcttgggagaaatgaatgaagatcacgt tcagaaagaatggaggaagagctgctcatttttgaagatgagggagaccaccact actatataatgaatagactaataccactgctcagtggttaacacacagcagtgattgc agctgccaatggaatgaagttatgggtgactgaaatataccaaacactgagggtg ggcaatggtgtctctgtg	8780	NM_003831 / NM_145906	SEQ ID NO: 45

SERP1 NB9	"serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9" serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9"	209723_at	+	ttgcaccattggccgtgtgtgtgaacctggccctcaagcaatccgccctacacag ccctcccaaaagtctagattacaggaataagccactgagccacagccctgtcagta tcctttatgaaattataaacatcigcaacattatgatacatatgcagatactatgcatctct ttattagtggtgaagtgctatgcaattatggtcttgaattccctcatcatgaattgca ttcaacacacactcttctgctcgttttatacatatgctttgcttataaagataattatccctct gtttatattctctcatctctgtattgcttttaa	5272	NM_004155	SEQ ID NO: 46
SIX1	sine oculis homeobox homolog 1 (Drosophila) / sine oculis homeobox homolog 1 (Drosophila)	205817_at		ccggaggcaagagagacggccggcgaggagcaaggaagggagagacaaccca aaacaataactctctccacaacagcagaaaccaactctctctctggaagggggca agccgtcatgtccagctcagaaggaattctccctccccaaggtccagaccag aactcgtctctctctgagggcaatattggccacgcagagagctcaactattctc tcocgggttaacagccctgcagccacgtccagctctgggacactggggtctaaagt ctccaaagactctctcggccctccacccagctctgggacactggggtctaaagt ggggaggggacggggccctcgaagggtattctctgagcagcaacacactgcagcact agggacactgtaaatagaaatcaggaacattttgcagctgtttctggaggtgtgcg cataaagggaatgggtggaacttcacaatatcttttaaaaaatcaaaacacacagcgat ctcaagcttaa	6495	NM_005982	SEQ ID NO: 47
SLOC3 A1	"solute carrier organic anion transporter family, member 3A1 / solute carrier organic anion transporter family, member 3A1"	219229_at		ggctgagccaccagtgagttcttggctctactctgacccctagacaacctggggaggga ccctgcccgaacacacagacacataaggcaaaagttatctataactgggaagcc atgagtggtgaaacacatgagctcgtttatagctataaaggagggctgaactct gtaataatccaaagggtatctttttcttaaaaaaagaaaaaaggttccaaaaaa accaaaactcagtaacacacacacaggaacagatgcacacacacagcagacagac acacgcactgtctctttctcagatcagagccagacagagattcagaataagga gaatgacatcgtcggcagggctcggagggccactcgcgcgggtgggccaacagag tctacttgaggccactcatgttttcaaggatgctgacagctgcaagcaacaggcact gccaaattcagggaacagtggtggccagctggaggtggac	28232	NM_013272	SEQ ID NO: 48
SPINK 1	"serine protease inhibitor, Kazal type 1 / serine protease inhibitor, Kazal type 1"	206239_s _at	+	gagacgtgtaagtgcggtgcagtttcaactcagccctggagcgcagaactcagcca tgaaggtaacacaggaactctctcagtgccctggtgcccctgtgagctctatcgttaacact ggagctgactccctgggaagagagcccaaatgttaacatgaacttaattgagtgac caagataatgacccctgctcgtgggactgaggaatactatcccaatgaatgcgtgt atgtttgaagggtcggaacccgaactctatctatcattcaaaaatctgggcccgtgctg agaacccaagggttgaaatcccatcagggtcacccg	6690	NM_003122	SEQ ID NO: 49

SPINK 5	"serine protease inhibitor, Kazal type 5 / serine protease inhibitor, Kazal type 5"	205185_at	+	agccatccatglttagagctctcaagaggaagacagccagacitcttcaagtctctg gattctgagatgtaacagactaccgagatgcccagagataggtctatcttggccaaa ggatttaagcctgctctgtgtgagatggccaaactatcaacaatccttgaatgctct gtcatgaaaactgatacccaaaaataacacacatccgcatgacaggaaggtgt gaggagagcaacccagccacccagccagccagatgcccgtctgacga atgacaggaagatgtgaaagccatgagggaataaataaacccagttctgaatc acctctccatccatctgtatatacaagaattctcgagctgtcttattgtatagaa aacaatacagagctttgggaatggaatcactgatttcaagcttcttccattctctcta gaatctgtatctgaggtataaagaacattccaccaagtgtgagccctcaaaatgccc tgattacaatgtctctgtccc	11005	NM_006846	SEQ ID NO:50
STC2	stanniocalcin 2	203438_at		gtccaatctctgaagcaatgattgacattggacaatcacaataaataaagcaaatg agtcatataaaataacccctctacttggctttatctgatacaacaatttactcatgagcc ttctttgaggaaagatgtggtatccaaataaagatttatttttttttttttttttttt ttaacaagatgattcgaacacccctctctgttgaatcaataatagccctgttattctgaagt gagaggaacagatagtagtaaatgctgacatctaaactaaataaataaataaataa ccagccagaaciatagtcatactcacacaaggaggaatttaaacctcgaaccaa gcaaaaggcttcacgggaataagcatggaaaacaatgcttccaggtgcccactctta aggaggacaacccctctgaatcagaattggcaccacgtgagctgtgctgaatgcat aatatctgttctactacggatttaggaacaggactgtacattgtccattgcat tgggggactttttgtgggacttaataaagattctttgatctaccgggaatatataatg tacagagtaacattggaatcattgtgaaaggaagcgaatgaaaggtcagagatgaa gtagcgaagtattggaatctctggaaggaatcactgtgtgaaatggaagagaca agtattagaccocaaagcaaacagcagagatgcaagagatgcccacaaa ggacaaggaacaattttctgtgacacttaacccggaagacitgtgtgtagaagaa aagaagctttgtgacactgtgaggagagaggggagggcagggcagatgcatgct gagcgtacagagcaagagcgtgctgtgctctccatccatgaaatgac taatttaccgtgaaggacccatgtgttatgttctctaatcttccatctccctaaagccctct gagagagatg	8614	NM_003714	SEQ ID NO:51
TAZ	transcriptional co-activator with PDZ-binding motif (TAZ)	202132_at		gcccgtgcgcacagatcgaggaaatgcatctccgggacgttggaacacacagacatga agataagaacccgttacggagctgtatctcaacccctgaaggatgccaagaacccgtg accgtgcgggaatgtgtgtggggccataacaccccaagcagatcgtgtgtaga ccctcagaagagatggcagtgatgagctgaaggatcctgaaggccatgaccaa ggaggccatccgaagacacagatggcccgactggcggaacgacagacacaccc gttccatcgcgcaatgtgaggaataaagaactgcacatcacacacaggtgcaagcc cgcatgctgtagagccatgacccactttgtgtgtgcaacgaggtgtggaacccgt ggaaagtcgtgacccctcgtgtagatgtgtgacgttgggcccctcccgcgcca cgctccgtgtgacacagctctctctgtgagacccctagaaaggcggtgcatgtccc	25937	NM_015472	SEQ ID NO:52
TCEA2	"transcription elongation factor A (SII), 2 / transcription elongation factor A (SII), 2"	203919_at			6919	NM_003195 / NM_198723	SEQ ID NO:53

TNNC1	"troponin C, slow / troponin C, slow"	209904_at	+	tgatgacatctacaaggctgggtagagagcagctgacagaaagagcagaaaaatga gtcaaggcagccttcgacatctcgtgctgggocctgaggatggctgcatcagcacc aaggagctgggcaaggtgatgagatgctgggccaagaacccccccttgaggag ctgcagagatgacatgaggtggagcagagacggcagcggcagcggcagccttg atgagctcctggctcatggtgctggcagagagcagcagcaaaagggaaatcig aggagctgctgacatctccgcatgttgacaaaaatgctgaggtacatcagcctg gatgagcagaaatgctgagcctcagcagcagcagcagcagcagcagcagcagc atcgaggagcctcatgaaggagcagcagcagcagcagcagcagcagcagcagc gatgagct	7134	NM_003280	SEQ ID NO:54
TRPM2	"transient receptor potential cation channel, subfamily M, member 2"	205708_s_at		accctggccatcagcggggcctgggctgagcctgggcccctggccagagt ccactccctcctggctgctcacccagagcagctcaccacatggagctcatggcc tgaggcaagttcccgaggagctgggtccctcctggcccccctcagcctatgctgt gaggaaaggccctgccactcccaagagagggccctcctgctgaggtgctcaca catggagcctggcctggcctggcctggcctggcctggcctggcctggcctggcct aaactccctggggtgacagtgagcagcagcagcagcagcagcagcagcagcagc agaggccctggcagggctggcccaaggacccctgggagcagggctgacagagc tcctccctcctcctccctgggagcagcctgctggccatgtggcagggagcagc gcaggagcggggagcgtggggcctcctggttggtgacacagc gaagccacaagaatgccacatgtagatatacagtagagagtgactccacagctc ctggagagcaatatagtgactgaagagtggggccccttgcttgctggatataag gggtgctctcctcctgtaattgggtggaaacactcggcttatggatcattaggctc tttcttaagagtagcttaaaatcaagagatccaatattttaaagccacaaagtagat acataatgacagagatttagcagtaaaatgttagaaatcaaacataaagaaatc aagctcttatttgctgggtatgctgattattttaaattccacccctccttattatca cttgtaagtgcccttgatgttgatgaaatgtagtagggagagtagcacaatgaaatg atggccctgctccctagctcctcctcctcctcctcctcctcctcctcctcctcct gaccttggttaatacaagaagtgatttgagccctccatcctcctcctcctcctcct gccgaagagcctgctgctggggtgctgcagcgcctcctggccctggcctcagc cgtggagctcagcggcagatcgagagcagagacactctggccatggctgcctga agaattctcctgcgaagatcccgctccacgtggctactgttccaaataccagaat acatcatggaaggtgtagaggccacattaaagtagcactcctcagcgaaggaatg tcccaaacctgtaacacctcctcctcctcctcctcctcctcctcctcctcctcct gacttggaacacactgcaaatattccatagcagcagataccacactcgtggttgccg atatgctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcct gcacatccagcggccgacccgttctcctcctcctcctcctcctcctcctcctcct gaggacatgagaaatcaggtgtgagcgcctcctcctcctcctcctcctcctcctcct gctgtgctc	7226	NM_00100118 8/ NM_003307	SEQ ID NO:55
VAMP4	vesicle-associated membrane protein 4	213480_at		tgagcacaagaatgacatgtagatatacagtagagagtgactccacagctc ctggagagcaatatagtgactgaagagtggggccccttgcttgctggatataag gggtgctctcctcctgtaattgggtggaaacactcggcttatggatcattaggctc tttcttaagagtagcttaaaatcaagagatccaatattttaaagccacaaagtagat acataatgacagagatttagcagtaaaatgttagaaatcaaacataaagaaatc aagctcttatttgctgggtatgctgattattttaaattccacccctccttattatca cttgtaagtgcccttgatgttgatgaaatgtagtagggagagtagcacaatgaaatg atggccctgctccctagctcctcctcctcctcctcctcctcctcctcctcctcct gaccttggttaatacaagaagtgatttgagccctccatcctcctcctcctcctcct gccgaagagcctgctgctggggtgctgcagcgcctcctggccctggcctcagc cgtggagctcagcggcagatcgagagcagagacactctggccatggctgcctga agaattctcctgcgaagatcccgctccacgtggctactgttccaaataccagaat acatcatggaaggtgtagaggccacattaaagtagcactcctcagcgaaggaatg tcccaaacctgtaacacctcctcctcctcctcctcctcctcctcctcctcctcct gacttggaacacactgcaaatattccatagcagcagataccacactcgtggttgccg atatgctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcct gcacatccagcggccgacccgttctcctcctcctcctcctcctcctcctcctcct gaggacatgagaaatcaggtgtgagcgcctcctcctcctcctcctcctcctcctcct gctgtgctc	8674	NM_003762 / NM_201994	SEQ ID NO:56
ZNF313	zinc finger protein 313	200868_s_at		tgagcacaagaatgacatgtagatatacagtagagagtgactccacagctc ctggagagcaatatagtgactgaagagtggggccccttgcttgctggatataag gggtgctctcctcctgtaattgggtggaaacactcggcttatggatcattaggctc tttcttaagagtagcttaaaatcaagagatccaatattttaaagccacaaagtagat acataatgacagagatttagcagtaaaatgttagaaatcaaacataaagaaatc aagctcttatttgctgggtatgctgattattttaaattccacccctccttattatca cttgtaagtgcccttgatgttgatgaaatgtagtagggagagtagcacaatgaaatg atggccctgctccctagctcctcctcctcctcctcctcctcctcctcctcctcct gaccttggttaatacaagaagtgatttgagccctccatcctcctcctcctcctcct gccgaagagcctgctgctggggtgctgcagcgcctcctggccctggcctcagc cgtggagctcagcggcagatcgagagcagagacactctggccatggctgcctga agaattctcctgcgaagatcccgctccacgtggctactgttccaaataccagaat acatcatggaaggtgtagaggccacattaaagtagcactcctcagcgaaggaatg tcccaaacctgtaacacctcctcctcctcctcctcctcctcctcctcctcctcct gacttggaacacactgcaaatattccatagcagcagataccacactcgtggttgccg atatgctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcctcct gcacatccagcggccgacccgttctcctcctcctcctcctcctcctcctcctcct gaggacatgagaaatcaggtgtgagcgcctcctcctcctcctcctcctcctcctcct gctgtgctc	55905	NM_016683	SEQ ID NO:57

Table 2: as described in priority application US60/619027 filed on 18/10/2004.

Gene Symbol	Gene Title	Affymetrix ID	+ up in sensitive	Sequence	Locus Link	RefSeq Transcript ID	SEQ ID NO.
AG:172L1	alanine-glyoxylate aminotransferase 2-like 1 / alanine-glyoxylate aminotransferase 2-like 1	221008_s_at	+	gctgaagaagccacacatgaactgcttagggacagcaccactgactccaaa gaaaatccacagcagagagagaatggaatggtgacggatcacacattcactgct cagtaagaggctcaagacacagactgattgcaitttaaaagcaagatgcgatgccc aggttacagagaatgagtagatggtctcctcgcgttaaatagctctattatcctct aaaagggaatgacgttagattacataaataaaaaggtaaaagaglaaactcaga ataaaccagagataaicaaacacatgtaagattattagtcagactagcctggt aatcttctagtgattctgaagctaccctgatttatttataaattgaagctgcaaa ctcaaatcaaatggcaattacctctcatgttttaattggtcaaatagagagca aagtaaacagggtccctcaccttttgagact gtgccatagtgcaggctgggagctttaagcctcagctcattataaocccagaaa acagagcctctatgaltacattctcgtacacatcctgacaggtacaaatcttataaattcaacta atgattgagggtccattatttagtggtactcagaaatggtcactttcctattacacgga gtgtgtaaaactaaagacatttgaacacacacagaaatgtctattgctatggga aatcttcttaaacaccagtgagggttagaaaagaaatgattatcttgtagcaaatia actttacatcttttctactgtatgtgtgttggtggaocgataaagtggttaactctga ggcaaaagtagaatagtttatgtaatagaagaaagaaatgtglaagttttga ttctactctatgtctggactgcattcacacatggcatgaataaagtcagggtcttta caaatgttatttgatagatactggaatgtgttgccatattggtccatt gggaltgatttggccattgtcaaaagtgcaagacacagcctctcttggccg acaaacttaaaaatccatgaagggtgctgcacacagatgagaaactctgacc aggaltgatttcccgagtgagattgacctgctcaacatccggagggaattc attgagaatattgacaagctctccacacagcattgagggtgacacctccgga gactcctgaaggcctgtgtgtctctgtgtgtggtgagactagggccacagctt tggcgggcatcttgcacagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagacacaaaggaagggtgaggggtggggggagggg gttgggtgggctctatctcatggagcttaggaacgctccacacccacgggc catcgaggggcagcagcgctgagcggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcaggaatgcaaatatataagacacctggagaca gtacatgcaaatgtcaagaaatatttctggtgaacggtgtgggaaaaagttcc atgaaacacacagacatgattgacagtagttatcaaaaattgcaatgcagcca tagctgctttatgtctgtgatactcaagctgtgtgttattacagtcacgttta gaagacaatcagcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgtaactgctatagcataactgaagataaaatta cagatatacaattggaactgcacagtcacagtcataagcaataaagatgagtggtgic ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaaaacttca attgtcacctttatgctatttctgata	64850	NM_031279	SEQ ID NO:58
AKAP12	A kinase (PRKA) anchor protein (gravin) 12 / A kinase (PRKA) anchor protein (gravin) 12	210517_s_at		gtgccatagtgcaggctgggagctttaagcctcagctcattataaocccagaaa acagagcctctatgaltacattctcgtacacatcctgacaggtacaaatcttataaattcaacta atgattgagggtccattatttagtggtactcagaaatggtcactttcctattacacgga gtgtgtaaaactaaagacatttgaacacacacagaaatgtctattgctatggga aatcttcttaaacaccagtgagggttagaaaagaaatgattatcttgtagcaaatia actttacatcttttctactgtatgtgtgtgttggtggaocgataaagtggttaactctga ggcaaaagtagaatagtttatgtaatagaagaaagaaatgtglaagttttga ttctactctatgtctggactgcattcacacatggcatgaataaagtcagggtcttta caaatgttatttgatagatactggaatgtgttgccatattggtccatt gggaltgatttggccattgtcaaaagtgcaagacacagcctctcttggccg acaaacttaaaaatccatgaagggtgctgcacacagatgagaaactctgacc aggaltgatttcccgagtgagattgacctgctcaacatccggagggaattc attgagaatattgacaagctctccacacagcattgagggtgacacctccgga gactcctgaaggcctgtgtgtctctgtgtgtggtgagactagggccacagctt tggcgggcatcttgcacagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagacacaaaggaagggtgaggggtggggggagggg gttgggtgggctctatctcatggagcttaggaacgctccacacccacgggc catcgaggggcagcagcgctgagcggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcaggaatgcaaatatataagacacctggagaca gtacatgcaaatgtcaagaaatatttctggtgaacggtgtgggaaaaagttcc atgaaacacacagacatgattgacagtagttatcaaaaattgcaatgcagcca tagctgctttatgtctgtgatactcaagctgtgtgttattacagtcacgttta gaagacaatcagcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgtaactgctatagcataactgaagataaaatta cagatatacaattggaactgcacagtcacagtcataagcaataaagatgagtggtgic ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaaaacttca attgtcacctttatgctatttctgata	9590	NM_005100 / NM_144497	SEQ ID NO:59
ANXA6	annexin A6 / annexin A6	200982_s_at		gtgccatagtgcaggctgggagctttaagcctcagctcattataaocccagaaa acagagcctctatgaltacattctcgtacacatcctgacaggtacaaatcttataaattcaacta atgattgagggtccattatttagtggtactcagaaatggtcactttcctattacacgga gtgtgtaaaactaaagacatttgaacacacacagaaatgtctattgctatggga aatcttcttaaacaccagtgagggttagaaaagaaatgattatcttgtagcaaatia actttacatcttttctactgtatgtgtgtgttggtggaocgataaagtggttaactctga ggcaaaagtagaatagtttatgtaatagaagaaagaaatgtglaagttttga ttctactctatgtctggactgcattcacacatggcatgaataaagtcagggtcttta caaatgttatttgatagatactggaatgtgttgccatattggtccatt gggaltgatttggccattgtcaaaagtgcaagacacagcctctcttggccg acaaacttaaaaatccatgaagggtgctgcacacagatgagaaactctgacc aggaltgatttcccgagtgagattgacctgctcaacatccggagggaattc attgagaatattgacaagctctccacacagcattgagggtgacacctccgga gactcctgaaggcctgtgtgtctctgtgtgtggtgagactagggccacagctt tggcgggcatcttgcacagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagacacaaaggaagggtgaggggtggggggagggg gttgggtgggctctatctcatggagcttaggaacgctccacacccacgggc catcgaggggcagcagcgctgagcggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcaggaatgcaaatatataagacacctggagaca gtacatgcaaatgtcaagaaatatttctggtgaacggtgtgggaaaaagttcc atgaaacacacagacatgattgacagtagttatcaaaaattgcaatgcagcca tagctgctttatgtctgtgatactcaagctgtgtgttattacagtcacgttta gaagacaatcagcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgtaactgctatagcataactgaagataaaatta cagatatacaattggaactgcacagtcacagtcataagcaataaagatgagtggtgic ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaaaacttca attgtcacctttatgctatttctgata	309	NM_001155 / NM_004033	SEQ ID NO:60
AREG	amphiregulin (schwannoma-derived growth factor) / amphiregulin (schwannoma-derived growth factor)	205239_at	+	gtgccatagtgcaggctgggagctttaagcctcagctcattataaocccagaaa acagagcctctatgaltacattctcgtacacatcctgacaggtacaaatcttataaattcaacta atgattgagggtccattatttagtggtactcagaaatggtcactttcctattacacgga gtgtgtaaaactaaagacatttgaacacacacagaaatgtctattgctatggga aatcttcttaaacaccagtgagggttagaaaagaaatgattatcttgtagcaaatia actttacatcttttctactgtatgtgtgtgttggtggaocgataaagtggttaactctga ggcaaaagtagaatagtttatgtaatagaagaaagaaatgtglaagttttga ttctactctatgtctggactgcattcacacatggcatgaataaagtcagggtcttta caaatgttatttgatagatactggaatgtgttgccatattggtccatt gggaltgatttggccattgtcaaaagtgcaagacacagcctctcttggccg acaaacttaaaaatccatgaagggtgctgcacacagatgagaaactctgacc aggaltgatttcccgagtgagattgacctgctcaacatccggagggaattc attgagaatattgacaagctctccacacagcattgagggtgacacctccgga gactcctgaaggcctgtgtgtctctgtgtgtggtgagactagggccacagctt tggcgggcatcttgcacagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagacacaaaggaagggtgaggggtggggggagggg gttgggtgggctctatctcatggagcttaggaacgctccacacccacgggc catcgaggggcagcagcgctgagcggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcaggaatgcaaatatataagacacctggagaca gtacatgcaaatgtcaagaaatatttctggtgaacggtgtgggaaaaagttcc atgaaacacacagacatgattgacagtagttatcaaaaattgcaatgcagcca tagctgctttatgtctgtgatactcaagctgtgtgttattacagtcacgttta gaagacaatcagcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgtaactgctatagcataactgaagataaaatta cagatatacaattggaactgcacagtcacagtcataagcaataaagatgagtggtgic ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaaaacttca attgtcacctttatgctatttctgata	374	NM_001657	SEQ ID NO:61

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BCL3	B-cell CLL/lymphoma 3 / B-cell CLL/lymphoma 3	204908_s_at	+	gggcagatcttgactatgaggggggggggggggggggggggggggggggg cttctgaaacgtggaagatctgacttgcgggggggggggggggggggggggg caggtattgcacagaaagacatgcactaccatacccccctctctctcaggac cctgttccccactcgtcctcccaaggactcgaacccagatctcaggcacc agtcctgtccggaatgccaccacatctcctcattccatgttccctccagagct ggggagaggg agctcaggacgctcgtcgaagggtggagcgcactggaagagaaacacagg tcttaagcgtgagaatcgcggacaagaglacacccagctccoaaggactcc agctccgctcgg gcaatgagatccaggaagcgtggcacatctggaagtcgctcgtcgtgtt tgccttgaaactcctctgactatcactcagctcaggggggtcatggggcaacgg ttagcgggagagacaggggtagcgggagagagggcctcggagcaggtcgtg gaggggcactggggcagctcgtgtgtgggacacagctcgggttgacccagg cctgtctcccaagagcctccctccggacaatgagtcctccctctgtctccac cctgagattgggcaggggggggggggggggggggggggggggggggggggg ttttttgggggggggggtgttttttgggggggggggggggggggggggggg cagctggacagatcaactcctcgtcggaccactggagagaaagaaatgacc acctccacgcccgcctccagagctcgtgagtcacacccggcagacagcct ggaatccagcagcagcgtcgggggggggggggggggggggggggggggggg ccaagagcccccacccgggacccacccctgcctcctgggtagggctcgtg ctgggactcaccctctggttagacactctcaagggtggtcctcaggggacc cctgtgtgtcgtcgtcgtggggcaccctcctgcctggggcctcccttggccta cctggggcagcccccacccctgcatgcccctgggggggggggggggggggg gaaccccccacctgctgcacccacacacacacacacacacacacacacac aggccttgagtgccacattaaatg	602	NM_005178	SEQ ID NO:63
BST2	bone marrow stromal cell antigen 2 / bone marrow stromal cell antigen 2	201641_at	+	gagctcaggacgctcgtcgaagggtggagcgcactggaagagaaacacagg tcttaagcgtgagaatcgcggacaagaglacacccagctccoaaggactcc agctccgctcgg gcaatgagatccaggaagcgtggcacatctggaagtcgctcgtcgtgtt tgccttgaaactcctctgactatcactcagctcaggggggtcatggggcaacgg ttagcgggagagacaggggtagcgggagagagggcctcggagcaggtcgtg gaggggcactggggcagctcgtgtgtgggacacagctcgggttgacccagg cctgtctcccaagagcctccctccggacaatgagtcctccctctgtctccac cctgagattgggcaggggggggggggggggggggggggggggggggggggg ttttttgggggggggggtgttttttgggggggggggggggggggggggggg cagctggacagatcaactcctcgtcggaccactggagagaaagaaatgacc acctccacgcccgcctccagagctcgtgagtcacacccggcagacagcct ggaatccagcagcagcgtcgggggggggggggggggggggggggggggggg ccaagagcccccacccgggacccacccctgcctcctgggtagggctcgtg ctgggactcaccctctggttagacactctcaagggtggtcctcaggggacc cctgtgtgtcgtcgtcgtggggcaccctcctgcctggggcctcccttggccta cctggggcagcccccacccctgcatgcccctgggggggggggggggggggg gaaccccccacctgctgcacccacacacacacacacacacacacacacac aggccttgagtgccacattaaatg	684	NM_004335	SEQ ID NO:64
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CAPZA2	"capping protein (actin filament) muscle Z-line, alpha 2"	201237_at		caatccacagatcaactcctcgtcggaccactggagagaaagaaatgacc acctccacgcccgcctccagagctcgtgagtcacacccggcagacagcct ggaatccagcagcagcgtcgggggggggggggggggggggggggggggggg ccaagagcccccacccgggacccacccctgcctcctgggtagggctcgtg ctgggactcaccctctggttagacactctcaagggtggtcctcaggggacc cctgtgtgtcgtcgtcgtggggcaccctcctgcctggggcctcccttggccta cctggggcagcccccacccctgcatgcccctgggggggggggggggggggg gaaccccccacctgctgcacccacacacacacacacacacacacacacac aggccttgagtgccacattaaatg	830	NM_006136	SEQ ID NO:66

CCND1	cyclin D1 (PRAD1: parathyroid adenomatosis 1)	208711_s_at		ggcggaggagaacaaacagatccatcccaaacacgcgcagacactctgtgoc ctctgcccacagatgaagttcaattccaatccgocctccatggcagcggg gagcgtggcgcagcagggcctgaacctgaggaagcccaacacactc ctgtctactacgcctacacgctctctccagagtgatcaagtgagccgga ctgctccggcctgacagagacatgaagccctgctggagcagcctgc gcaggccacagacacatgaccacacagccgcgaggaaggagaaaga ggagagagaggggagcctgctgacacacacacagcgcggacgt ggacatctgagggccacagggcgccgcacccgcacccgcacccgcagag ggcggagccggccacagctgctccctgacagcctccctccgagcatttg ataccagaggggaagctcattctctctgtgtgtgtgtgtgtgtgtgtgt ctccatctcagacttaagcaaaa	595	NM_001758 / NM_053056	SEQ ID NO:67
CCND1	cyclin D1 (PRAD1: parathyroid adenomatosis 1) / cyclin D1 (PRAD1: parathyroid adenomatosis 1)	208712_at		gtttgggtatgttaattctgactgctgtctgtgtgtgtgtgtgtgtgtgt ataatgtaattaaagagactccaaatctcaatgaagccacagctcacgtgt gtgcccgtctatcagcagctgcgaaacaaagaatgcatccocgcgtgc ggccacagctgtggggccctgcccggcaggggctcctgtctcgaggcc atctcggcacagggccaccccccgcacccctccagaaacacggctcagct acctcaactctgtgtgctgctgtgtgtgtgtgtgtgtgtgtgtgtgtgt acgt agaagccaaagcgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt gacgcgaagctgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt aatctcgcattctggggtctgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt ctctgt ggatgacacccgggacacgtttatctatgaagaagagggcgagagaa gaggaacaggt tgaagt ccgcctgccatccagatgaatggaatgtgtgtgtgtgtgtgtgtgtgtgt ggctgt aggagcgt caaagaccaggt tggctgacatgacggagggcg	595	NM_001758 / NM_053056	SEQ ID NO:68
CDH1	"cadherin 1, type 1, E- cadherin (epithelial)"	201130_s_at		ctctgt aatctcgcattctggggtctgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt ctctgt ggatgacacccgggacacgtttatctatgaagaagagggcgagagaa gaggaacaggt tgaagt ccgcctgccatccagatgaatggaatgtgtgtgtgtgtgtgtgtgtgtgt ggctgt aggagcgt caaagaccaggt tggctgacatgacggagggcg	999	NM_004360	SEQ ID NO:69
CDKN2A	"cyclin-dependent kinase inhibitor 2A (melanoma, p16, inhibits CDK4) / cyclin-dependent kinase inhibitor 2A (melanoma, p16, inhibits CDK4)"	207039_at	+	cttttcaactgt gt catttgaactagggaagctcaggggtgtgtgtgtgtgtgtgtgtgtgtgt tagcaaatggca	1029	NM_000077 / NM_058195 / NM_058197	SEQ ID NO:70

CDKN2A	"cyclin-dependent kinase inhibitor 2A (melanoma, p16, Inhibits CDK4)"	209644_x_at	+	1029	NM_000077 / NM_058195 / NM_058197	SEQ ID NO:71
CEACAM 7	carcinoembryonic antigen-related cell adhesion molecule 7	206199_at	+	1087	NM_006890	SEQ ID NO:72
CHORDC 1	"cysteine and histidine-rich domain (CHORD)-containing, zinc binding protein 1 / cysteine and histidine-rich domain (CHORD)-containing, zinc binding protein 1"	218566_s_at		26973	NM_012124	SEQ ID NO:73
CLU	"clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J)"	208791_at	+	1191	NM_001831 / NM_203339	SEQ ID NO:74

CLU	"clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J) / clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J)"	208792_s_at	+	agcagcgaacgagaggaatttaacgggtgltccggcggtgcaaacctcacgcaa ggcgaagaccagtatactgcgggtgcaacggtggttcacacactctgact cggagcttcctccgtgtcactgaggtggtggaagctcttgactatgacct cactgacgggtccctgtagaagctccaggagaaaccttaattatggagac cgtggcggaagaagcgctgcaggataaccgcaaaagcaacgggagaggt gagatggtgagtgtgc	1191	NM_001831 / NM_203339	SEQ ID NO:75
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CTSB	cathepsin B	200838_at		tcccccgtgactagtgccgtgggagtaacctgcgccacgctgctggccccc ccgtgatccatccatccagggaggaagacagagacgaggaiggaagc ggagtcciaacaggaatgaagttcccccatactgctcccccagtaacctcaagc aagtagcttccacattgacacagaatacagag	1508	NM_001908 / NM_147780 / NM_147781 / NM_147782 / NM_147783	SEQ ID NO:78
CTSB	cathepsin B / cathepsin B	200839_s_at		tgggtgggagcccttggagaacgccagctctccaggtcccccgtcactatcga gtttgcaatgcaacaacctctgactctgtctcagcatgattctttaataagattt attttctgcactctcctaatcatgtgggtgagccagtggaacagcgggagcctg tgtgtgttgcagatgctctcctaatgacggtgctcaaaaggaaacacagtggtc aggagttgtctgaccacactgactctactaccacaaagggaataatgtaggaga aacacagctttactgttt	1508	NM_001908 / NM_147780 / NM_147781 / NM_147782 / NM_147783	SEQ ID NO:79

DAB2	"disabled homolog 2, mitogen-responsive phosphoprotein (Drosophila)"	201278_at		ggaacgttccagttcaattcagtcctgttgagcacagttctgaagggtttatf t'gtcaaaalaagttttgtttgtttttatgtgggttttaattgtgtcttgcoccttaa tgcctggtctgtggaggttaalccagcacatccaangttacctgagggggaa gaagggtgtatgtctcaagaagctaaacagagggccacalgcacctctca t'gtatagccccaagtagaaagctcctgtgtttatgttaaatggtaatggtacat alatggcataattttctacagctctctacagctacacalaaacacagactgaat agttcttaaatgtccaaatacctaagtgtctaaactggaggtaactattctagg tagttgaattttgaaagtcattgatcagccacaacactgttttgcataact	1601	NIM_001343	SEQ ID NO:80
DAB2	"disabled homolog 2, mitogen-responsive phosphoprotein (Drosophila)"	201280_s_at		aatccttattgttcaagttgtttgtgggtttcgtttcagagcataaaacctaaaggtt alatgtagaacaagccacctctttaaagaatactgttcacagacatcagttaca gagaattcttaagaiaaaattgaagcaactcaacactctcttagacactttgga atctaacccttaaggacacttttaagaagatagatccttcttctgaagatcaatt ctccaaaggccaagattgtcctttctccattctgtcagctatgcaaatgaggg aagaacattatcctcctccctttttctgtactttttcagctgtttgtctcctg ggttcaagtagtattaccacctttccaagccaacagactc	1601	NIM_001343	SEQ ID NO:81
DIAPH2	diaphanous homolog 2 (Drosophila) / diaphanous homolog 2 (Drosophila)	205726_at	+	gctcacctacattcattgcacacaaatgaattttcaccttttaagatgcattctgg tgcataaacacagatcgaaagttgtctnaaagctatgtcgcagggtcgtgc atgctctgtgttaattggatggacaggtctattcctaaatttgggtgatactttgctact atgggcatttaactgaaaataataatcgtatcccaactc'tgtcctgtatgtaacct ctctgccttttatgacacctttgacaaaatgcctctatgtgtccacagtcagcc acaaaactac'tc'gatacagaaggttctttaacagctttatcacatcagtgaaat ccctccacctaaaggagaggtgaaagccaagactccttgaatgggtattgag ggagattgttccatacccaagccacctgaaagaa'gtattcacttgcagtagaac t'gtggatttgc'tgcatttccaccttggaaataaacacctatc'taagcaggacca a	1730	NIM_006729/ NIM_007309	SEQ ID NO:82
EMP1	epithelial membrane protein 1 / epithelial membrane protein 1	201324_at		caccaaattaccatggd'gaggttagagagattggcagcaaaaactgtggga agatgaactgttcattatgatttcaatcacaatgatfatagaaggc'tc'ttagtgc aaaaaacatactacattcagacataccaaggggaatactcacattt'gtitaaag aag'tgaacctalgc'ggaggaacctat'gattccctatcttttactttttc'tg'gac atttat'gtc'at'g'aatttgcattactc'gg'gtatt'gtc'c'ag'act'g'att'ggg'ctctt cg'ttaat	2012	NIM_001423	SEQ ID NO:83
EMP1	epithelial membrane protein 1	201325_s_at		ttatcgccct'gagaagatctaccaggaggagaatct'gagacatct'goc'acttctc tttat'gactt'ctccatc'acattcttttatacccttctttt'gggag'tg'at'gcaat'g atttt'gt'attat'g'aaaa'gattat'ctaat'ct'at'ct'at'gttatt'clagt'aag gaaat'gt'gag'ggag'ccacc'aaatt'acctag	2012	NIM_001423	SEQ ID NO:84

EMP1	epithelial membrane protein 1	213895_at		aaggactggatcttctctgaggaalaaaggaatggataaagacatgataatcctg tgcnnnnnccagancnalaacaataaaggagggttttaattggaagcaggcaat ctnccagccctctggtcttgatgaatagttgcacagagattgacccaana atacacaatggaggctgaagaatgtaacataatatttaagtaataaacaattgca ttgattctgagcttctctagaaggctctcatgattctagattgctctgataaactc ataagggttccacnccctcattttagctcccccagggaattcttccccatgca tacaccagctcctaaatacaaccccaaggctatctctccatccctctgcagagg gaactttgcaagacatgcacaacaacctctagctctatccaagctgctctgctg ctaagattggatctctctcccaaaagccctggatgggaatgggggggaattgac agaattctcc	2012	NM_001423	SEQ ID NO:85
EREG	epiregulin / epiregulin	205767_at	+	taaaacctgactgaaccacttgaattttgctccaatataccatctctgagacttt gaaaaaaagtttttaattgatgccaatatattctgaccgttaaaaaaatctctgic atattggagaaggggaggtaattgactgtacaacacagatttctgggtatattta atgttttaaaaaagagtaattcatttaaatatctgtattcaaatgtatgattaaat gtaataatgattttctttttttttttgcaactgtaattgcacttttaagttgaagac caatttggtaaacggtttttaaaagatgctatggaacataaaagttgattgcatgca atttaagtaacttattgactatgaattattcggattactgaattgatacaattgttt gtttcaatatacagcttgataattgtgtaacctaaag gggactcatttagctctagaccactaatcaaaagttcggcatgagctcatgat ctatgctgtttctatgctggaagcaccggatgggggtatgagcaaatctgccc tgctcagcagtcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactggaatcactaactgactgaatgtaattggaatggcaataagct ttgtcccaagatgctggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgtg atattctctcaggctctgacaggctctctggaacctccacatatttttaactgca gtataaagtcagaaaaataaagttaacataacttccadlaacacacacatatgag atttcacaaaatccactataattggcaaatggttgaataataatttttagtaatt gcaatgcaaaattttctagctccactcttctccctgctttcttttttggggagctg gtaactgatgaatctttccaccctttctctcaggaataataagtggtttgttgg taacgtgatacatctgataatgaacactggagggaacacatctactgaattct gtaatttaaaatattttgctgctgataactatgaacagataagaatacttaacagat gctgdataaataaaggaataataaatttcaatcaataaaatgctatttttaaat ctattctctataattgatttctaatcaatgaattgactcttatttcttat	2069	NM_001432	SEQ ID NO:86
FGF2	fibroblast growth factor 2 (basic)	204421_s_at	+	gggactcatttagctctagaccactaatcaaaagttcggcatgagctcatgat ctatgctgtttctatgctggaagcaccggatgggggtatgagcaaatctgccc tgctcagcagtcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactggaatcactaactgactgaatgtaattggaatggcaataagct ttgtcccaagatgctggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgtg atattctctcaggctctgacaggctctctggaacctccacatatttttaactgca gtataaagtcagaaaaataaagttaacataacttccadlaacacacacatatgag atttcacaaaatccactataattggcaaatggttgaataataatttttagtaatt gcaatgcaaaattttctagctccactcttctccctgctttcttttttggggagctg gtaactgatgaatctttccaccctttctctcaggaataataagtggtttgttgg taacgtgatacatctgataatgaacactggagggaacacatctactgaattct gtaatttaaaatattttgctgctgataactatgaacagataagaatacttaacagat gctgdataaataaaggaataataaatttcaatcaataaaatgctatttttaaat ctattctctataattgatttctaatcaatgaattgactcttatttcttat	2247	NM_002006	SEQ ID NO:87
FGF2	fibroblast growth factor 2 (basic) / fibroblast growth factor 2 (basic)	204422_s_at	+	gggactcatttagctctagaccactaatcaaaagttcggcatgagctcatgat ctatgctgtttctatgctggaagcaccggatgggggtatgagcaaatctgccc tgctcagcagtcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactggaatcactaactgactgaatgtaattggaatggcaataagct ttgtcccaagatgctggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgtg atattctctcaggctctgacaggctctctggaacctccacatatttttaactgca gtataaagtcagaaaaataaagttaacataacttccadlaacacacacatatgag atttcacaaaatccactataattggcaaatggttgaataataatttttagtaatt gcaatgcaaaattttctagctccactcttctccctgctttcttttttggggagctg gtaactgatgaatctttccaccctttctctcaggaataataagtggtttgttgg taacgtgatacatctgataatgaacactggagggaacacatctactgaattct gtaatttaaaatattttgctgctgataactatgaacagataagaatacttaacagat gctgdataaataaaggaataataaatttcaatcaataaaatgctatttttaaat ctattctctataattgatttctaatcaatgaattgactcttatttcttat	2247	NM_002006	SEQ ID NO:88

[illegible]

ITGB2	"Integrin, beta 2 (antigen CD18 (p95), lymphocyte function-associated antigen 1; macrophage antigen 1 (mac-1) beta subunit) / integrin, beta 2 (antigen CD18 (p95), lymphocyte function-associated antigen 1; macrophage antigen 1 (mac-1) beta subunit)"	202803_s_at			atctggaggctctgatccacctgagcgacctccgggagglacagcgcgctttgag aaggagaagctcaagcccccagtggaacaatgataatcccttttcaagagcgc caccacgacgtgcatgaaccccacagtgctgagagtaggagcactggtgaa gacaggccgtgcaagcccccacacagtgctgcccacacgcggcgagacatg gctggccacagctctgaggaiglcaccaataacacagaaatccagttattcc gcccataaatgacagccatggccggccgtgcttctgggggctcgtcggggg gacagctccactctgacggcacagcttgagggagactgagggaggggtga gggtggtaggttaggtgctggttctctgtgcaagt	3689	NIM_000211	SEQ ID NO:94
KHDRBS 3	"KH domain containing, RNA binding, signal transduction associated 3 / KH domain containing, RNA binding, signal transduction associated 3"	209781_s_at			cagcccgccaggtggagtgtagtaccacgagggagcgccaactccagagg agctctgtccaccggggccagtgagtcggggagagggagactctcactccca gagcaagagagtgcccccacacgtgagtcggggagagggagactctcactccca cacaagagactatggagaatagactatgagtgatagtgatagtgacgtgctatgat gaacagagtgatgactctatgaiaacagctatgacccccagccccaaagtgtg gctgattactatgattacggacatggactcagtgaggagactatgattctacgg gcaagaagtgaggactaaactcaagaacacaaaggacactcaagggagacagc aaaggcgtctacagagacccagccatagggagactgattgactctgctgag tttgaaatagccaatctccaccagctctgtatctg	10656	NIM_006558	SEQ ID NO:95
KRT13	keratin 13 / keratin 13	207935_s_at	+		gagaacacggggcagagagcaggagtcgcgtatgcctgcagcgcagcag atccaggactctacagcagcagtcaggccacgctgagcgagctccgagtg agatggagtgccagaaacaaagagacagatgctgctggacatcaagacacg tcggagcagggagatgcacactaccgacgctcgcggaggcgacgacgccc aagaaagcgcagcccccagacactctgtaacacgactctagtcctctgta ccaccactctaatgctctgctgcgcgacactcgaigtcgagggcctaaatc gctggcgtccctccctctgctcagcccccagagggagagagagcgggca gtccctgcagagagagaggggctgctggacccaaaggctcagctcctgct cicaggacccccgtcctgactctcctgagtgggggccctctgtgctctctcc ggcggatctctcctctcgaactggatagcgttgggttctcaactc aaccttaacttagagctcattatgaagaatgaaacacacactctgagttgatt cccaaaagttaaaagcccttaagctcactgatttcaactcttgcacata gtcattaccctccacagccgttggtgcatagaggggtggtggttgattgat tttttaactctgagtgagaaatagataggggacaaaacactctggttcttaag acaatcagtgctgagcactctgacgaatggaatgaaatcactgtagccaat tagaatatttatgattgattggttggctgattttatgaaataataattatcatt ctgactctcgggaagcaa	3860	NIM_002274 / NIM_153490	SEQ ID NO:96
LASS6	LAG1 longevity assurance homolog 6 (S. cerevisiae)	212446_s_at	+			25378 2	NIM_203463	SEQ ID NO:97

LTBP2	latent transforming growth factor beta binding protein 2 / latent transforming growth factor beta binding protein 2	204682_at	<p>gggagccaaggccttatacgtctaaagaaatattcagtagctgaatcgcgcca glatagcctgtgggcacacgaagcaagggtgcctgggatacagcaacca tctacaagccctctattatacaacacgtctctacaggaaacaaacctctctg ggatctcctttgtgaaacacagtttgatgtctaaagataaaaagctatttccag tgtggctgtgtcagaagcagcaatttccaatgtgttttccctccaccagaa accctgcccttcctcagaaacacgatgcaggcattcctgagtttacaagc agagactcaccccaaacatagcggg</p>	4053	NM_000428 / NM_032035	SEQ ID NO:98
MAP4K5	mitogen-activated protein kinase kinase 5	203552_at	<p>acacacatgcaatttgcctaaacaaagattttataatacacagtttatacagaa ttacctaaagggaagctctatgttttcaacacagatagttgtaaggatcctaca gaagatattgatgattgaaataattctagaaaggggtgtatgtctagctgtgic taacatgtgatgtattctgcaacgacgataaaatccctgtattttcttacaata gggalaatgcataaggaattatctcatatattatcatccctaataatglacaggg ggaagttaattgccatgccatgatatgtattttactatactatgccagaggaact ataaagaattacacatgtaactctgtgggtttcacatattaggtattcatttgagta gggtgaagagaanaaaaalatttaaatgaaatgaaattcctgagggatagta aat</p>	1183	NM_006575 / NM_198794	SEQ ID NO:99
MAP4K5	mitogen-activated protein kinase kinase 5	203553_s_at	<p>gaactctgcattctcatgttttaca gaaattgttgcaggcagccagcagtagatt ccattctgltaacacagttggagagagataacggttttagtgttttagacaacattgt gaaaattgtaaatctacaaggaaaattaaatcaagtaagaaactgacctctga gttaagttttgatttgcattgaaactgtgattgattgccttcacagacagtgthggcttct tggaaacatgggagtgagggtgaaaaggtcaagtcagatgaggtttaccaggga gattccagatgaaacaagatttccgttattaggatcagacagaggggtgctgttttg gaaagtagggcaacagaaaatctctacgcaacacgaaatctctacatctggct ggacatgaaaatagttctaa gcaaca gaaactgactcctaaatgcacaggaaa atgaatatctactcattgaaagggaataaggaattcaatacaaacatgcacta tgaattgctttaact</p>	1183	NM_006575 / NM_198794	SEQ ID NO:100
MMP2	"matrix metalloproteinase 2 (gelatinase A, 72kDa gelatinase, 72kDa type IV collagenase) / matrix metalloproteinase 2 (gelatinase A, 72kDa gelatinase, 72kDa type IV collagenase)"	201069_at	<p>ctcagagccaccctaaagagatccttgatattttcaacgcagccctgctttggg ctgcctgggtgtgccacacttcaggctctctctcttcaacaactctgtggctcac agaacacttgagccaalggagactgtctcaaggagggcactgtgtggcccgac agccctggccacagggtgcggacaggcattggccagggtggccactccagac ccctggcttcaactgcgtgccttgtaagaaactttctacattgaagtttgctgtat gcactgtttttcttgggtctgttttttttccactgaataatgcattctgcagaca aggacacaggtgtctgaagctcactgcacagtgcaatcagccacacatagtatg gttccctgtcactctactagcaatgtccctaccaggactctctccactggatgga ggaaaacacagccgtgctccctccgctcagccctccctgcccctcccttcaacact tcccatttgggaaat</p>	4313	NM_004530	SEQ ID NO:101

MYC	v-myc myelocytomatosis (avian) / v-myc myelocytomatosis viral oncogene homolog (avian)	202431_s_at	+	+	SEQ ID NO:102
NRP1	neuropilin 1	210510_s_at			SEQ ID NO:103
NRP1	neuropilin 1	212298_at			SEQ ID NO:104
OLFM1	olfactomedin 1 / olfactomedin 1	205591_at	+		SEQ ID NO:105
OLFM1	olfactomedin 1	213131_at	+		SEQ ID NO:106

OSMR	oncostatin M receptor/ oncostatin M receptor	205729_at	+	9180	NM_003999	SEQ ID NO:107
PCDHAC 2	"protocadherin alpha subfamily C, 2/ protocadherin alpha subfamily C, 2"	210674_s_at		56134	NM_018899/ NM_031883	SEQ ID NO:108
PDGFRA	"platelet-derived growth factor receptor, alpha polypeptide / platelet- derived growth factor receptor, alpha polypeptide"	203131_at		5156	NM_006206	SEQ ID NO:109
PDZK1	PDZ domain containing 1 / PDZ domain containing 1	205380_at		5174	NM_002614	SEQ ID NO:110

PEA15	phosphoprotein enriched in astrocytes 15/ phosphoprotein enriched in astrocytes 15	200788_s_at	taaattcaatgcagctcagagactatttagacaaagttcaagttaggagacttita ggatgggagtaaaacttaattggaggggagggtgctgctggaagaag gaagaagccagactggttagacagctacttaactcctagccagcctacgagc cctggccctctggccactgctgagacacactgcttaacacacacacacactagg actccacagtttgccttaaggaactcccaagctcccttcccttctgctgctc ccttaagaagagagagactactttagaattgggtgggggaalagacatgaac tgtcctccattgggatatgttacttagagtgagagagataaaggagccttcc ttatggaagaatgggagagagagacaggggtcttccagagagctagtag ttctctgtaaggcaaaataatctaaagagactaacctgcccacccactccttat tgctgtgagattgcccc	8682	NM_003768 / NM_013287	SEQ ID NO:111
PGM1	phosphoglucumutase 1/ phosphoglucumutase 1	201968_s_at	cggaccatccaaagtcactgattgaagagcatgacagaacaaaatgtatca ccaaagcatttagatttgccttttcaacacagttgacgagcagtgcatttaca ggcactgccaaacaagaatgccctgggagcgtggaggaaagagaccctggc ggcttagatcaatcactcaattccttctatgcctcctgcatgctgctgggtatt gtccttagccatcaggtacagtttaccactacaatgtaagctatagggtgagcat cagcagtgagtgaggccactctctactagatggatggcgaatgaaatgatggg caagttccttctttgtgaatcttccccacttctgttttaccatgtaacccaa aatgcaattctagtgctctctgccaalcagttcttctctgagtgagacgactg gclacagattctgctgttttggacattgic	5236	NM_002633	SEQ ID NO:112
PI3	"protease inhibitor 3, skin- derived (SKALP)"	203691_at	gattggtatggccttagctcttagccaaacactctctgacaccatgagggccag cagctcttgatcgtgggtgctctcctcactcgtggacgctggtcttagagcagc tgccacggagttctgttaaaaggtcaagacactgtcaaaaggcctgttccattca atggacaagatccctgttaaaaggacaagtttcagtttaaggtcaagataaagtca aagcgaagaagccagtcacaaggctccactcactaagccctgctcctgcccc attatctgacccgtgctgctgctgaatccccctaaacccctgcttgaagaatact gactgccacaggaatcaagaggtgctggaaggctcttgcgggaggtgctgttgcg ttcccaagtgaaaggagccggctcctgct	5266	NM_002638	SEQ ID NO:113
PLAU	"plasminogen activator, urokinase / plasminogen activator, urokinase"	205479_s_at	cccgaccgtgggcatgttgaggcccatggtgagaatgaataattcccaatt aggaaagttaagcagctgaggtctcttgagggttagccaatgtgggagca gggtttgggagcagagacaciacgactcaggcagggtcctgatatcca tgaatgtracaggaataataatgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt gagtgaaagtgtgagtaagagcgtgctgctgagtgtaagtaataattccttaaa ctgtgtggactgtgatgccacacagaggtgtcttcttgagagaggttataggctc ctggggcctctgggtccccacagtgacagtgccgggaatgtacttattctgcag catgacctgtgaccagcactgtctcagttcacttccactatagatgtcccttctggc cagttatcccttcttagcctgactgtcatcaaatcactcactggtgtgggg	5328	NM_002658	SEQ ID NO:114

PLAU	"plasminogen activator, urokinase / plasminogen activator, urokinase"	211688_s_at		5328	NM_002658	SEQ ID NO:115
PP1F	peptidylprolyl isomerase F (cyclophilin F) / peptidylprolyl isomerase F (cyclophilin F)	201489_at		10105	NM_005729	SEQ ID NO:116
PP1F	peptidylprolyl isomerase F (cyclophilin F)	201490_s_at		10105	NM_005729	SEQ ID NO:117
PTGS2	prostaglandin- endoperoxide synthase 2 (prostaglandin G/H synthase and cyclooxygenase) / prostaglandin- endoperoxide synthase 2 (prostaglandin G/H synthase and cyclooxygenase)	204748_at	+	5743	NM_000963	SEQ ID NO:118

[illegible]

SPRY1	"sprouty homolog 1, antagonist of FGF signalling (Drosophila)"	212558_at		10252	NM_005841 / NM_199327	SEQ ID NO:124
SPRY2	sprouty homolog 2 (Drosophila) / sprouty homolog 2 (Drosophila)	204011_at		10253	NM_005842	SEQ ID NO:125
SFIPUL	sushi-repeat protein / sushi-repeat protein	205499_at		27286	NM_014467	SEQ ID NO:126
TCF8	transcription factor 8 (represses interleukin 2 expression) / transcription factor 8 (represses interleukin 2 expression) / transcription factor 8 (represses interleukin 2 expression)	208078_s_at	+	6935	NM_030751	SEQ ID NO:127
TGFA	"transforming growth factor, alpha / transforming growth factor, alpha"	205016_at		7039	NM_003236	SEQ ID NO:128

TGFB2	"transforming growth factor, beta receptor II (70/80kDa) / transforming growth factor, beta receptor II (70/80kDa)"	208944_at		glttgatggtgaaggtctcatattttagatttttaagatacatgcaaaaggtttgg aaatagaacctttagaacccctctcaagtggtgggtgggtgagagthtaagaca gltggtgtcagtagcatagagcgccctagaaatccactgcaacctgagggca tgcgataccatcccaatagctgtgcccattgacclaggtggtgagttctagaat acgtgccatcaicagatattcaagattcaagagattctcaactctgggtatcag cataaactgggaatgtagtgcaggagatctggtggt	7048	NM_003242	SEQ ID NO:129
TIEG	TGFB inducible early growth response / TGFB inducible early growth response	202393_s_at		tttgcctgcaagttctgtgagattgaaatgtaataccaaatgtgtttctgtagactct aagatcacatgcacttgttagaaaaaaacacgaagatgaatataatattgtaa agaaggatataaagaactcttagataaactctgaaaaagatggcttatgtcalca gtaaagtaacttattgtatgagatataatgtgtcttattgaatgaataaattgt gacattattcaaggttggaacaatgtgtctgttaattatagaggttttgggga tgtgaggtggtgggtagaaaaaattatagaacattcaattgtgtaaacgatttct ctttattctgttatatggtgatgatacacagtggaacaaacaaaatgacattgct taaaatataatgtaaaaaatgtaactatattccatttaacattgttttgtatattgg gtgagattctgacatcaaaacttgacccttggaacaaacaaagttaataaaa aaaaactctgtgactcaaatgtgac	7071	NM_005655	SEQ ID NO:130
TIMP3	"tissue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201147_s_at		gagtcggagatgagcagcacacacaaattcccagcccagtgatgctgtgt tgaccaagttgtcgtgagtcgtgagcaagcaccagccagcaataaacaagct ttctagtgtggaagactaaacatctgcctgaggtcagaggaacaaattgcctgcc ttgtacaaaagctcaggtgaaagactgagatgaatgtcttctcctcctgcctccc accagactctcctggaacacgtgttgagattggccagagactttttatgt aaattggataaatacacacacacatcacatccacagatatagccaagtagattt gggtagaggatactattccagataatggtttagctcacttagggggatattgtgt atatcaattgcatataccacacagggg	7078	NM_000362	SEQ ID NO:131
TIMP3	"tissue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201148_s_at		ttgtgtcgtgtgtgttgagaaatcatgacattccaagtgcacttttttttcatitt aattaaaattgaaattgaaacccgtcagcaacctctctccatcatgggtcalt ctgaccctctcctcctctgtcctgtctcatgtttggggccttcttaactgcctt cctgtgttagctcagatggcagatgaagtgatgtaacagggcctgggcaag aggaagactgcagagttgcctgcctgcctggctggaggaacactctcctgg gtggagacagctgtgtcctctccctagcctcctgtgtgtggaatgcaactcc tgagatctcactcctctggaaataaaattgtgtcactggggaaagcctgagttg caaccagttg	7078	NM_000362	SEQ ID NO:132
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TIMP3	"tissue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201150_s_at		gacatttttgaaatagccctgtctagggccaaactgtggccccagggagacacaccc ctccatgccacagacctgtctgcatgagacaatgacaatctggactacccc aagatggccaccaggtgttggctctgctacacaaaggttaacatgtcactagag tatattatagagagacaaacattatataaactcagtcgcaaaagcaaaacaaat ggaaagtaggggggtgtgagacacactccaaatggctctggaggc gagagaaaggagagactggagaaatgtttgtttggggtagaggtctt agattccacagactcccttccctttagccagctgtctgtccgaacocagaa gtgtgagagaaacacaaagagatcgaacccctgttagaaggatglat ttgtgtcctaaattctgtagccactgtttacagtttccatctgtattatg gaglatctacagagcttgcacccctccattttgtcttggctgtcttaattggct aatgcaccccccacaaagaaatcaccacaaataacttaataatgcaccaa aaggcaagactgacctagaaatctagccgtttggagatactaaactgtctca ggaaagactgttggatgtcatgcatgaacccatgttgcaatcaaatgataa aatagtattctattttcccccaccccgaaatgtcaaaatgtccatgtaaac ctgtacaaatggcagcttatcatalgcaatgtgaataatcatctgtgattgg aatgtctgtcatcccccaagtttcaagatttaagattctctctactactcia cgtttaatatcttgaagttgtattaaatgtatttaagattctctactactcia tgtaaatgtaaacgtgaagatgataaacgtgaagcagataactggaaccac ciaaagactccattttggagatttttgccttggctgtgttggaattat accaggctctcatgaatccacacttaactcgaacacgtggcaataaattia tctgtgtgactgtgataatataatcacactattgtgtgactgacacactgac aagtaaaaggcttgttcgaagatgtgtcaatgaagcacaatagatgaga tcaagatgacatgtccagacacagacagacagaaaggtcaactgtctgt aatgtgacactcatggaagaaagagcgtatgacacattgataaagat ctcaaaaagccaactcttctgtctgtcagagaaatcagatcatctcaca ggacattactgtgactcagaataatcaactcagaatgaatcacaagctt ggctcagagtaaaacacaaatcagttctgagatatagcaattgattigan aagattcttaagctgtgtaatactgtctgtgtt	7078	NM_000362	SEQ ID NO:134
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TNFSF10	"tumor necrosis factor (ligand) superfamily, member 10 / tumor necrosis factor (ligand) superfamily, member 10"	202688_at	+	gtagcagctcacataactgggacacagaggagagaaacacacatgtctctcc aacctcagaatgaaaggctctggccgcaaaataaactcctgggaatcat caaggagtggtggtcattctctgagcaactgcaactgaggaatggatgactggt catccatgaaaaagggtttactacatctatcccaacacacttctgattcagga ggaaataaaagaaacacaaagaaacagacacaaacaaatgtccaatatattac aaatacacaagatctcctgacctatgtgtgagaaaggtgtgagaaatgtgt tggtctaaagatgcaaatatggactctatccatctcaaggggggaatattga gcttaaggaaatgacagaattttgtctgtaacaaatgagcaactgataacat ggacacatgaagcaggttttgcggcctttttaggtggcactgacat ctctacatatacagttgtctgagaaatcagagactgtcagcttccaaaca ttaatgcaatgttgaacatctgtcttataatctactctgtgaagactgtgaa gaaagcacaacatctctcagagtagratcagtagtaggtccaggt tcttaaggagacacatctctaaagaaagagagagaggcaccacataaaa gactgcaggttgcctgggtgcagtggc	8743	NM_003810	SEQ ID NO:138

TUBB4	"tubulin, beta, 4"	213476_x_at	tumor suppressor candidate 3	209227_at	10381	NM_006086	SEQ ID NO:139
TUSC3	tumor suppressor candidate 3	209227_at	tumor suppressor candidate 3	209227_at	7991	NM_006765 / NM_178234	SEQ ID NO:140
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TUSC3	tumor suppressor candidate 3	213423_x_at	tumor suppressor candidate 3	213423_x_at	7991	NM_006765 / NM_178234	SEQ ID NO:142

Example 4**RT-PCR Confirmation Studies**

In addition, the sequence of the RT-PCR primers used in the confirmatory follow up studies as highlighted in Figs 3, 4, 5 and 6 are listed in Table 3. Note that DAPK2 was not identified by Affymetrix analysis, only via follow up of the DAPK gene family by RT-PCR following discovery of predictivity of DAPK1. Hence no Affymetrix ID or Affymetrix ID sequence is provided for DAPK2.

Table 3

Sequences relevant to genes followed up by RT-PCR (see Figs 3, 4, 5 & 6)

(all sequences written 5'-3')

Gene	affy id	affy probe seq	Tagman Forward Primer	Tagman Reverse Primer	Tagman probe
EMP1	201324_at	CACCAAATTACCTAGGCTGAGGTTAGAGAGATTGGCCAGCAAA AACTGTGGGAAGATGAACCTTTGTCATTATGATTTTCATTATCAC ATGATTATAGAAGGCTGTCTTAGTGCAAAAAACATACTTACATT TCAGACATATCCAAAGGGAATACTCACATTTTGTAAAGAAATT GAACATGACTGGAGTAAACCATGTATTCCTTATCTTTTACTT TTTTCTGTGACATTTATGTCCTCATGTAATTTGCATTACTCTG GTGGATTGTTCTAGTACTGTATTGGGCTTCTTCGTTAAT	AGCCATCCTG CCCTTCTGA	ACCTTACAAAC TCTCTTTCC	CAAAGCA AAACATC ACATTCC AGTC
NES	218678_at	GCAGCACTCTTAACCTACGATCTCTTGACATACGGTTTCTGGC TGAGAGGCCCTGGCCCGCTAAGGTGAAAAGGGGTGTGGGCAA AGGAGCCTACTCCAAGAATGGAGGCTGTAGGAATATAACCTC CCACCCCTGCAAAGGGAATCTCTTGCTGCTCCATTCTCATAGG CTAAGTCAGCTGAATCCCGATAGTACTAGGTCCCTTCCCTCC GCATCCCGTCAGCTGGAAAAGGCCGTGTGGGCCAGAGGCTTC TCCAAAGGGAGGGTGACATGCTGGCTTTTGTGCCAAGCTCA CCAGCCCTGCGCCACCTCACTGCAAGTAGTGACCATCTCAC TGCAGTAGCACGCCCTCCTGGGCCGTCTGGCCTGTGGCTAAT GGAGGTGACGGCACTCCCATTTGTGCTGACTCCGCCCATCCCT GCCACGCTGTGGCCCTGCCTGGCTAGTCCCTGCCTGAATAAA G	GCCCCCTTCA GGAGGAGGA	AGTGCCGGGG AGATGGTCTT	AGTGCTC TGAAGAC CTCTTGG GC
DAPK1	203139_at	CCTCCTCCAGGGTGATTTTATGATCAGTGTGTTGCTCTAGGA AGACATTTTCCGTTTGCTTTTGTTCGAATGTCAATGGTGAACG TCCACATGAAACCTACACACTGTCATGCTTCATCAATCCCTCTC ATCTCAGGTAGAAGGTTGACACAGTTGTAAGGGTTACAGAGAC CTATGTAAGAATTCAGAAGACCCCTGACTCATCATTTGTGGCA GTCCTTATAATTGGTGATAGCCAGATGGTTTCCACATTTAG ATCCTGTTTTCATAACTTCCTGTACTTGAAGTCAAAAGCAGAA AATAAAGGAAGCAAGTTTTCTTCCATGATTTTAAATTGTGATC GAGTTTTAAATTGATAGGAGGGAACATGTCCTAATTCTTCTGT CCTGAGAA	AGGAAACGCT ACCTCTCTGT	CTGGAGGAGG ATTCCCTTCT	CTTGCTG TATGCTG ATCATCG CC
DAPK2	Not applicable	Not applicable	GGGTAGGCAC CTGGCATC	AGTGCAGTGG CGTGATCTC	TACTCCA GGGGCT GAGGTGA CA

Example 5

Diagnostic test for Clinical Studies

The predictive gene lists above have been generated using the preclinical studies described. The following approach is employed to develop a diagnostic test for the clinical setting based on this data.

- a) Identify patients which represent the population of individuals whom we would expect to derive benefit from a diagnostic test, and for which pre-treatment tumour samples and outcome of gefitinib treatment are known or will be available. For each sample the expression level for our genes of interest is evaluated, using for example the RNA signal from RT-PCR. QC procedures are applied to identify the set of samples and genes to take forward to step b).
- a) Identify a subset of the genes which together are able to distinguish between patients showing different responses to gefitinib. There are a variety of methods which are useful to select the subset of genes and combine their expression values to provide a prediction, possibly a predictive value and a corresponding threshold which distinguishes between different patient groups. An example is stepwise Linear Discriminant Analysis where genes that distinguish well between patient groups are successively added to a linear combination until addition of a further gene does not provide additional predictive power (Mardia et al.). The threshold value of the linear combination is then selected to give the appropriate sensitivity and specificity properties.
- d) Tool validation would partly be carried out during development in step 2, for example using cross validation and permutation tests. In addition, the finally developed diagnostic procedure (gene subset and method of combining to generate a prediction and a platform for biological analysis) is tested and validated in its entirety using an independent set of samples not used within tool development in step b).

References

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CLAIMS

1. A method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or DAPK2, whereby to predict an increased likelihood of response to the erbB receptor drug.
2. A method according to claim 1 comprising testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7, DAPK1, ACOX2, GSPT2, TNNC1 or DAPK2.
3. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7 or DAPK1.
4. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of at least two of NPAS2, NES, CHST7 or DAPK1.
5. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of at least three of NPAS2, NES, CHST7 or DAPK1.
6. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of NPAS2, NES, CHST7 and DAPK1.
7. A method according to any preceding claims additionally comprising testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein.
8. A method according to claim 7 comprising testing a biological sample from the mammal for expression of any one of EMP1, SLC20A1, SPRY2 or PGM1.
9. A method according to any one of claims 7-8 comprising testing a biological sample from the mammal for expression of EMP1.

10. A method according to any preceding claim wherein the tumour is selected from the group consisting of leukaemia, multiple myeloma, lymphoma, bile duct, bone, bladder, brain, CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head, neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural membrane, peritoneal membrane, prostate, renal, skin, testicular, thyroid, uterine and vulval.
11. A method according to claim 10 wherein the tumour is selected from one of non-small cell lung, pancreatic, head or neck.
12. A method according to any preceding claim wherein the erbB receptor drug is selected from any one of gefitinib, erlotinib, PKI-166, EKB-569, HKI-272, lapatinib, canertinib, AEE788, XL647, BMS 5599626, cetuximab, matuzumab, panitumumab, MR1-1, IMC-11F8 or EGFR11.
13. A method according to claim 12 wherein the erbB receptor drug is gefitinib.
14. A method according to any preceding claim wherein the mammal is a human and in which the method comprises testing a biological sample from the human for increased expression of DAPK1 and decreased expression of NPAS2, NES, CHST7 and EMP1 whereby to predict an increased likelihood of response to gefitinib.

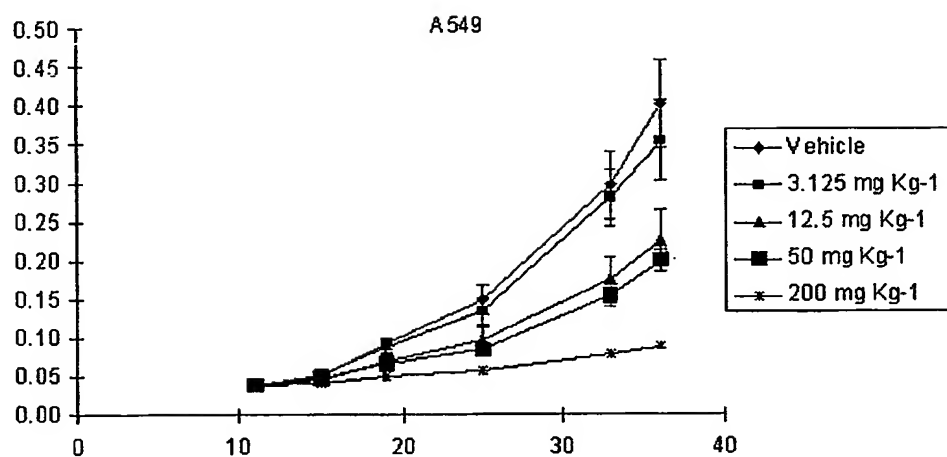
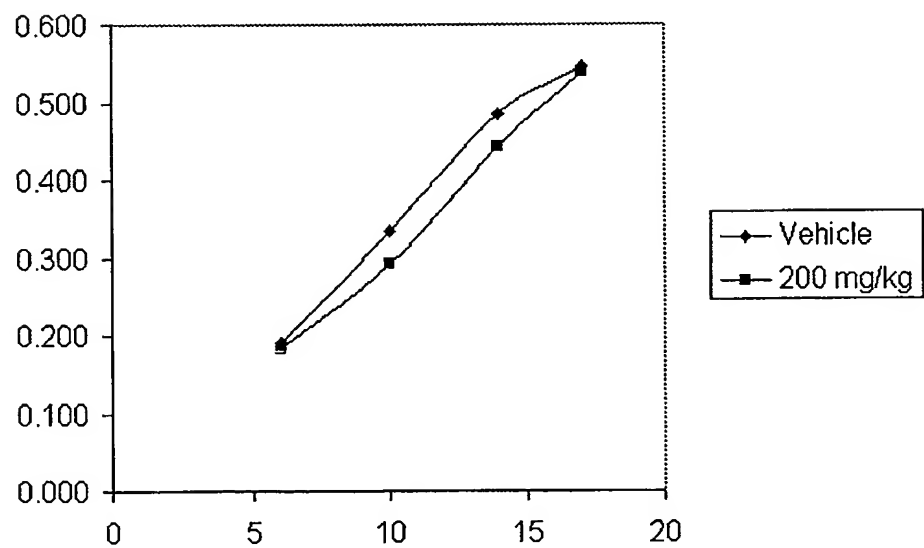
Figure 1**Figure 2**

Figure 3

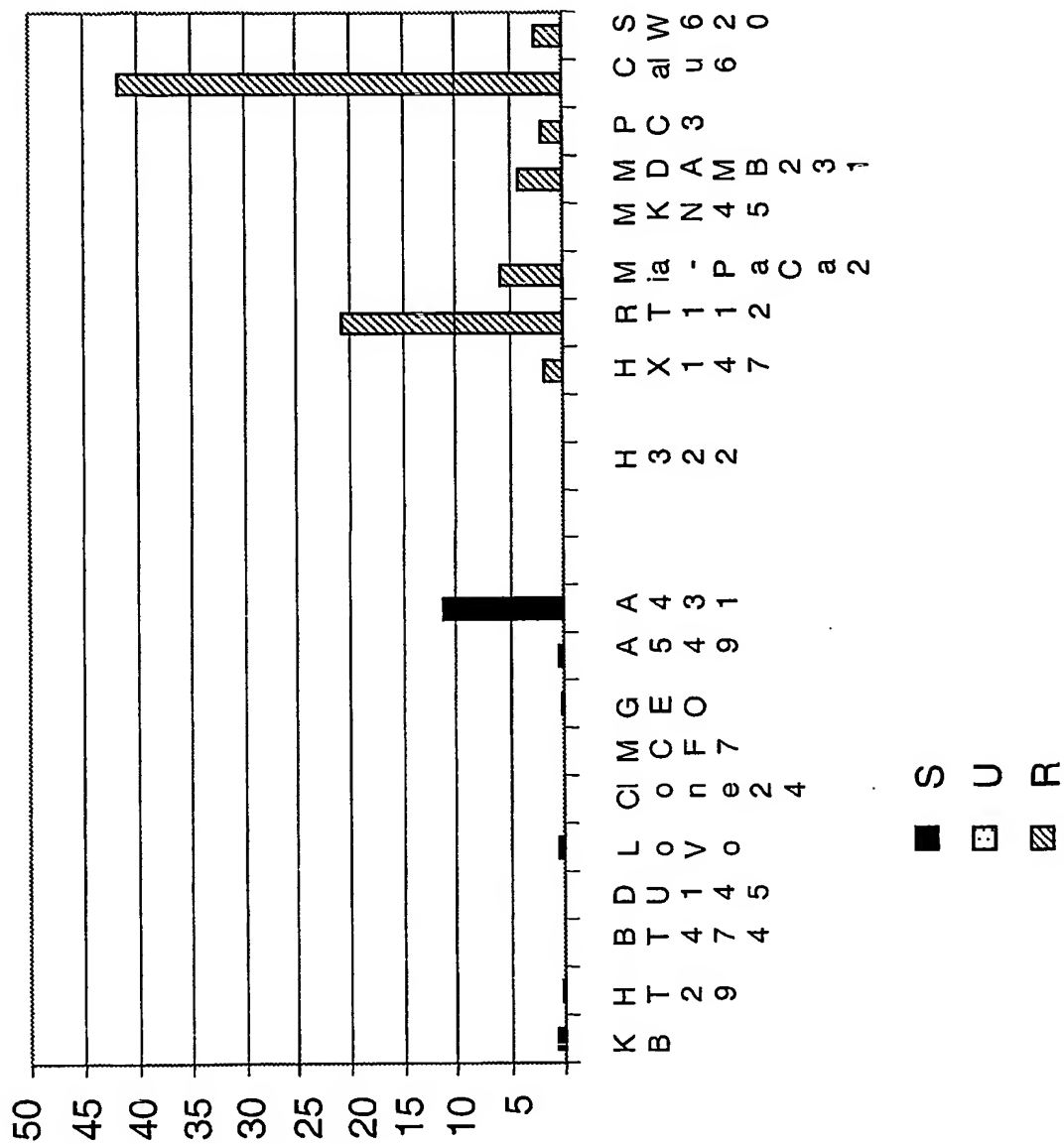


Figure 4

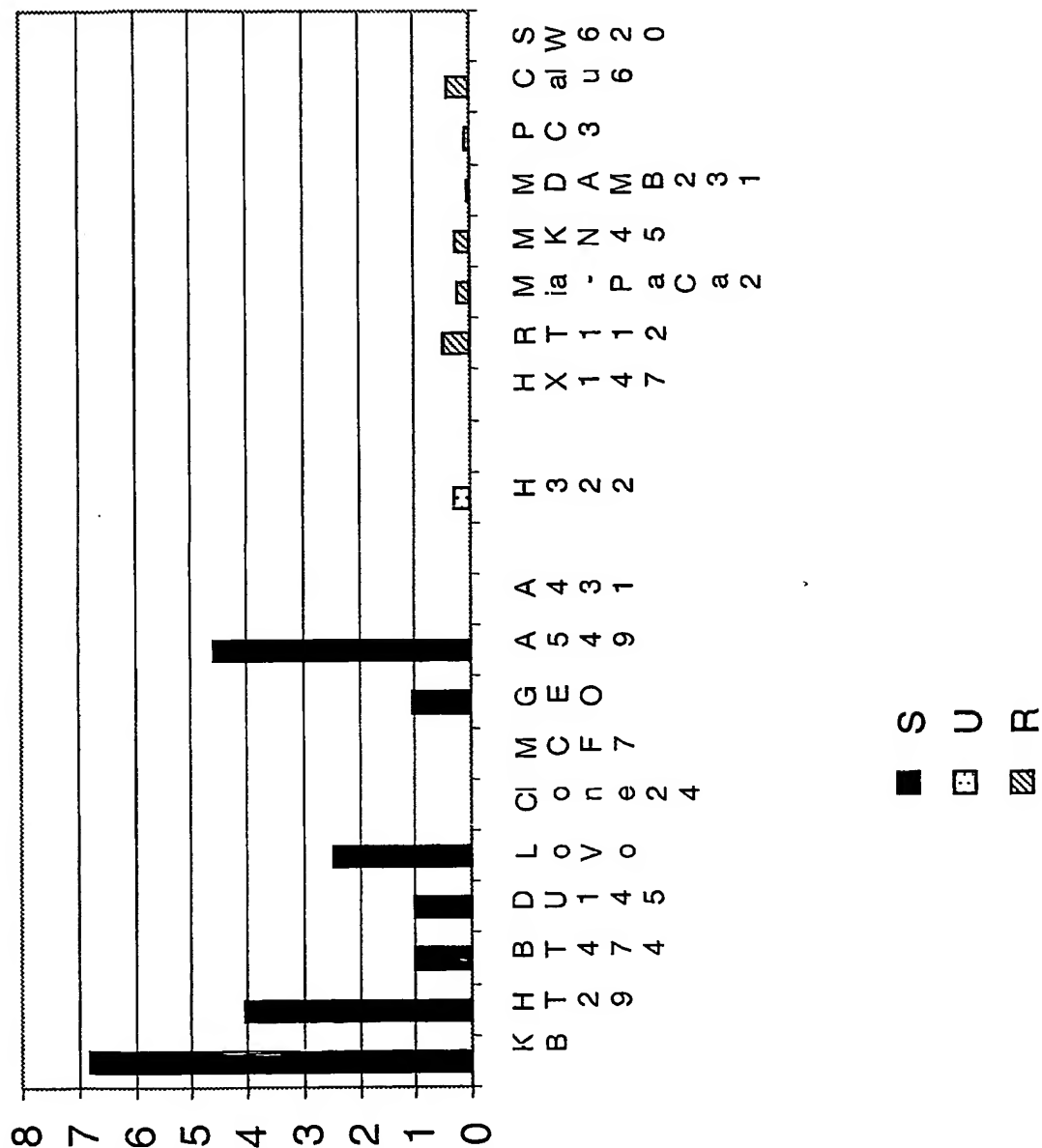


Figure 5

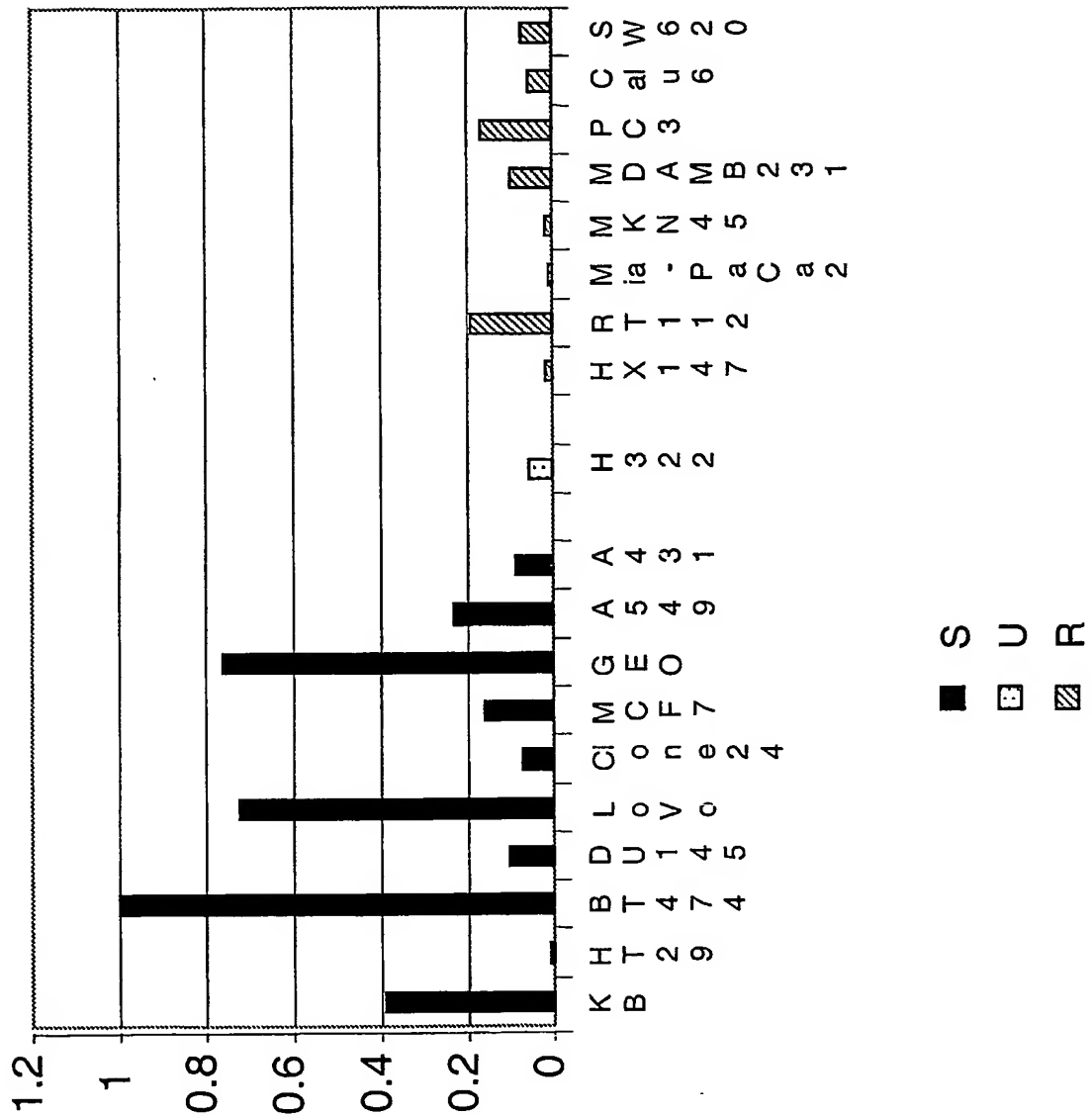
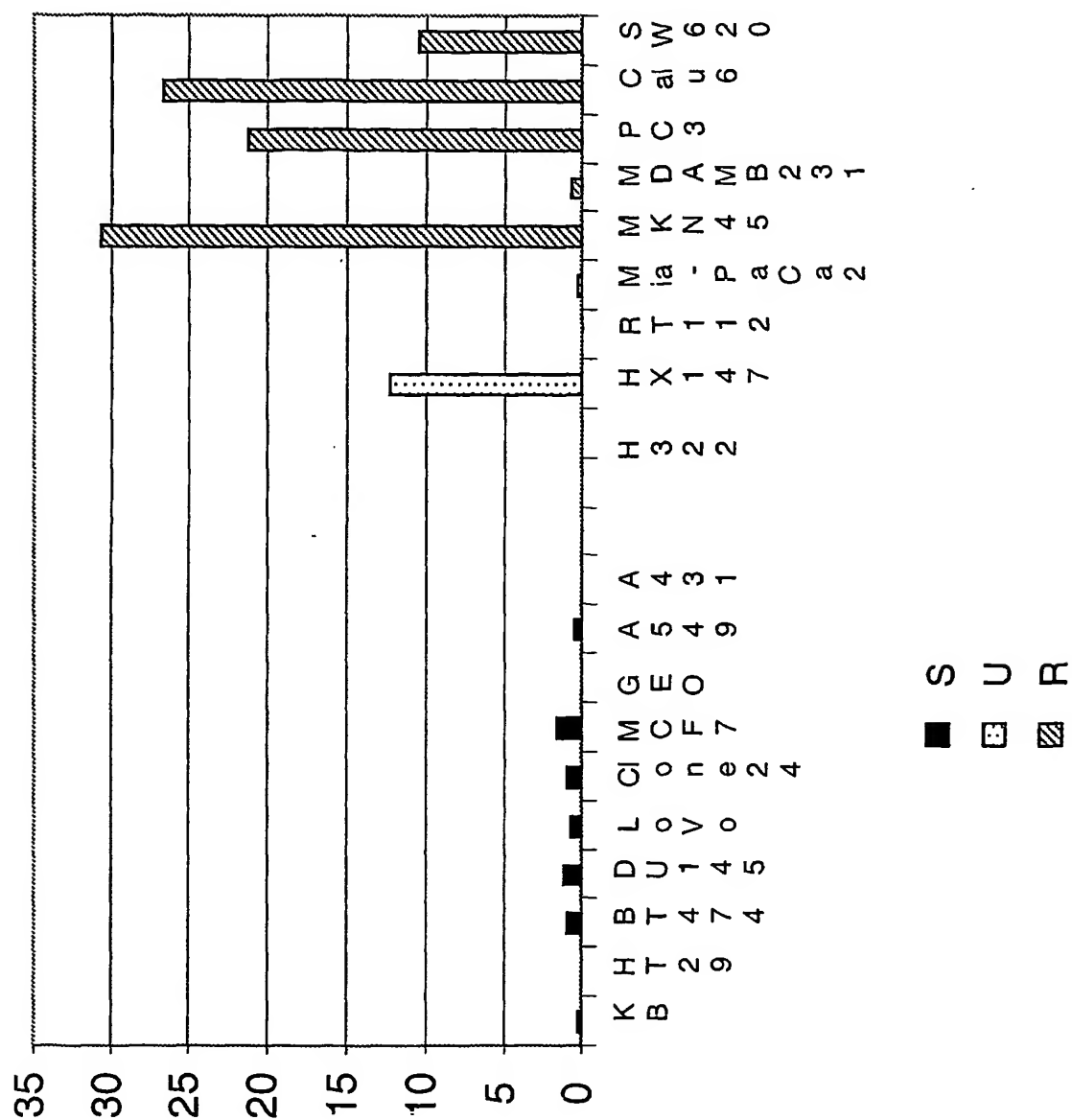


Figure 6



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tcaacagtcg aagaagggtgt gggcagaaga aaaagctagt gatcaacagt ggcaatggag 300
ctgtggagga cagaaagcca agtggactca acggagaggc cagcaagtct caggaaatgg 360
tgcatttggt gaacaaggag tcgtcagaaa ctccagacca gt 402

<210> 8
<211> 417
<212> DNA
<213> Homo Sapiens

<400> 8
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aaaagagaca aaagacatct tcgaatccat atttcaagcc tggtagaatt ggcttttcta 120
gcgaacctt tccaaaagt ttatattgag attcataaca acaccaagaa ttgattttgt 180
agccaacatt cattcaatac tgttatatca gaggagtagg agagaggaaa catttgactt 240
atctggaaaa gcaaaatgta ctttaagaata agaataacat ggtccattca cctttatggt 300
atagatatgt ctttgtgtaa atcatttggt ttgagtttcc aaagaatagc ccattgttca 360
ttcttgtgct gtacaatgac cactgttatt gttactttga cttttcagag cacaccc 417

<210> 9
<211> 546
<212> DNA
<213> Homo Sapiens

<220>
<221> misc_feature
<222> (104)..(104)
<223> n is a, c, g, or t

<400> 9
ttctatgcat ccacacaaa atcctgcaga atgtaagtaa gctctgcttt ataagatggg 60
ttcaccttca tcgcagactg aaagtttcag tttttatttt tttncagaaa gcacgaaaaa 120
ttatttataa tagtctggag aaaaaacaca ctgtaatat tcaagtgtat gcagtagaat 180
gtactgtaac tgagcccttt ccacatgctc taggctocaa tgtctcctgt aggtccacct 240
aactgtgtgt tttcaggagc aatgccatcc atgtttgtgc tgtagacttg ctgctgctga 300
atcctttctg gggactttct catcgggcag ggagcagagg gcttctcggt catgcacct 360
ttgcctgaac acctatgtag ctgctgtggt gtgtatatat tactcttaag aggagtgtgt 420
gtgtctgtgt ttgttttaaa agtcacttat ttcttacagt gatttcaatt gcaccatgac 480
ttcttacta aaaccacaaa gtctgtctta aaactatgga aaacctaacc tgattagagc 540
cttgac 546

<210> 10
 <211> 546
 <212> DNA
 <213> Homo Sapiens

<400> 10
 ggcaatctgt cacactctca gagtctggga cttgacttgc taccaacaac tgctgtgcaa 60
 ttctgctgag caggaatatc atgagctgtt caataatgac ggacgcattg gttgagatga 120
 agtttccagt aaggaagtga cagtgcattg tggatatatta tggctgtaaa ataggaagag 180
 ctttagttcc caggctgaac ctgccactgc tggagccatt tcaacaaggc atcctcacia 240
 caaagaagag atgtgatttg gtaccatttc acaccagcag gtgtctggac gaaaacatca 300
 atgtgaataa gggccaagtg cagtcctgtc ttgattaaat tacttaataa tattattaaa 360
 taataatagg tctgggcagt attgttttta acctgactca tccagctgtc cttcaaatag 420
 ctccgtctcc ctctaccag aactgatttt taaaaagaag taatttttct ccctgggctg 480
 ggaaaaccct aatgaactga aacacacttt tactttaaaa ttttctgtc tggcgttttt 540
 gtaatc 546

<210> 11
 <211> 496
 <212> DNA
 <213> Homo Sapiens

<400> 11
 gaattcccta gaaatcctac tgggaagtat aggcagatct ctccctcata taacggatgt 60
 ttcttggcgc ttggaatatc agataaagac caatcaactt cataggatgt acagacctgc 120
 atatttggtg accttaagtg tacagaacac tgattcccca tcctatccag agattagttt 180
 tagttgcagc atggaacaat tacaggactt ggtggggaaa cttaaagatg cttcgaaaag 240
 cctggaaaga gcaactcagt tgtaacttgg ggaagttaac gatccgcccg agtgcagagg 300
 aaaaccagaa acgccttgcc ttcagctgaa ccaccgtttg tgcgagctgg atgtcctttt 360
 cagtagaaaa gaattttcct tttgaattta taccattcat caattttgac actttaaaaa 420
 cgtgtgaaag ggtaagagg gaaagatact gcccaagtat ttgaatcgtt tagtagtaac 480
 tgtccattta tcctat 496

<210> 12
 <211> 313
 <212> DNA
 <213> Homo Sapiens

<220>
 <221> misc_feature
 <222> (190)..(190)
 <223> n is a, c, g, or t

<400> 12
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 caataattac ttagtatcat ctaatatgtg gttcatattt aaatttggtg ttttgagatg 120
 ggtcttacaa ttggtttatt caattgcatt ttttctaact cgtgtctcaa gtgttttaaa 180

5

aatctaactgn acttataatg acttatataa tgtattttctc attttacctt tttccaaaa 240
gaggaaataa tggcaaacca tataatatgtg tacatttcact gtcaaaaagc aaacccttgt 300
tttgataact tgt 313

<210> 13
<211> 395
<212> DNA
<213> Homo Sapiens

<400> 13
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cttttgttcc aatgtcaatg tgaacgtcca catgaaacct acacactgtc atgcttcac 120
attccctctc atctcaggta gaagggtgac acagttgtag gggtacagag acctatgtaa 180
gaattcagaa gaccctgac tcacatttg tggcagtcct ttataattgg tgcatagcag 240
atgggttcca catttagatc ctgggttcat aacttcctgt acttgaagtc taaaagcaga 300
aaataaagga agcaagtttt cttccatgat tttaaattgt gatcgagttt taaattgata 360
ggagggaaca tgcctaatt cttctgtcct gagaa 395

<210> 14
<211> 569
<212> DNA
<213> Homo Sapiens

<400> 14
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gttctgaagt ttacagatg gaacctgaca atgtgaatgc cctgaaagat cgagctgagg 120
cctatttgat agaggaaatg tatgatgaag ctattcagga ttatgaaact gctcaggaac 180
acaatgaaaa tgatcagcag attcgagaag gtctagagaa agcacaaaga ttattgaaac 240
agtcgcagaa acgagattat tataaaatct tgggagtaaa aagaaatgcc aaaaagcaag 300
aaattattaa agcataccga aaattagcac tgcagtggca cccagataac ttccagaatg 360
aagaagaaaa gaaaaaagct gagaaaaagt tcattgatat agcagctgct aaagaagtcc 420
tctctgatcc agaaatgaga aagaagtttg acgacggaga agatcctttg gatgcagaga 480
gccagcaagg agggggcggc aaccttttcc acagaagctg gaactcatgg caaggggttca 540
atcccttcag ctcaggcgga ccatttaga 569

<210> 15
<211> 481
<212> DNA
<213> Homo Sapiens

<400> 15
tgagggccac gggcttggtt agtggaagg gtgtttggga aattgttaaa tcagttaccc 60
gtagtagagc tatttcttgt acttctaagt tttctagaag tggaaggatt gtagtcatcc 120
tgaaaatggg ttacttcaa aatccctcag cctgtttctt cactactgtc tatactgaga 180
gtgtcatgtt tccacaaagg gctgacacct gagcctggat tttactcat cctgagaag 240
ccctttccag taggggtggc aattcccaac ttccttgcca caagcttccc aggtttctc 300
ccctggaaaa ctccagcttg agtcccagat acactcatgg gctgccctgg gcagccagca 360

6

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ttcattgtaa gttccctctt tgaaaactgg tgtgtgggtg ttcagttctg tgtctggtgg 420
gtatggacag acagtaatct cctgtgatct gtgctagctg tgaggcagct ctggaacctg 480
a 481

```

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<210> 16
<211> 398
<212> DNA
<213> Homo Sapiens

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<400> 16
ggctcccgagc aagggtagga cgggccgcat gcgggcagaa agttgggact gagcagctgg 60
gagcaggcga ccgagctcct tccccatcat ttctccttgg ccaacgacga ggccagccag 120
aatggcaata aggactccga atacataata aaagcaaaca gaacactcca acttagagca 180
ataacggctg ccgcagcagc caggggaagac cttggtttgg tttatgtgtc agtttcactt 240
ttccgataga aatttcttac ctcatTTTTT taagcagtaa ggcttgaagt gatgaaaccc 300
acagatccta gcaaagtgtc ccaaccagct ttactaaagg gggaggaagg gagggcaaag 360
ggatgagaag acaagtttcc cagaagtgcc tggttctg 398

```

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<210> 17
<211> 499
<212> DNA
<213> Homo Sapiens

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<400> 17
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tatgtgccgc cgagcctgcg cgacggggcc agccgcgcgc gggagtccat gcagcccaac 180
cgcagagccg acgacaacgc caccatccgt gtcaccaact tgtcagagga cagcggtgag 240
accgacctgc aggagctctt ccggccttcc ggctccatct cccgcatcta cctggctaag 300
gacaagacca ctggccaatc caagggtttt gccttcatca gcttccaccg ccgagaggat 360
gctgcgcgtg ccattgccgg ggtgtccggc tttggctacg accacctcat cctcaacgtc 420
gagtgggcca agccgtccac caactaagcc agctgccact gtgtactcgg tccgggaccc 480
ttggcgacag aagacagcc 499

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<210> 18
<211> 261
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (41)..(42)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (196)..(196)
<223> n is a, c, g, or t

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<400> 18
atgtgtcggg gagagagccc gcagggaagg gtaaagccca nnggggcagg gccctccag 60

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atgcctgagg aggggggcagg tccccctccc tctcctcctc tccccctccc atctaaaggg 120
 gtttgggggag agacacaggc aggcgagggg gctggtcccc agtctgttgg ggtggtgctc 180
 agggtaaaagg gctatnggca acagggggacc agaccaggga tgagtgggga gggcacaagg 240
 accatttgcc agaatccacc g 261

<210> 19
 <211> 526
 <212> DNA
 <213> Homo Sapiens

<400> 19
 ctgttgctcc aggatgcatt ctgataggag ggggcggcag ggctgggcct tgtgacaatc 60
 tgcctttcac cacatggcct tgcctcgggtg gccctgactg tcagggaggg ccagggaggc 120
 agagcgggag ggagtctcag gaggaggcct gccctgaggg gctggggagg ggttacctca 180
 tgaggaccag ggtggagcct gagaagagga ggaggtgggg gcttgagggt gcttggtagc 240
 tgagggggacg ggcaagttag aggggagggga gggaagtcct gggaggatcc tgagctgctg 300
 ttgcagtcta acccactaat cagttcttag attcagggga agggcaggca ccaacaactc 360
 agaatggggg ctttcgggga gggcgcctag tccccccagc tctaagcagc caggagggac 420
 ctgcatctaa gcatctgggt tgccatggca atggcatgcc cccagctac tgtatgcccc 480
 cgacccccgc agaggcagaa tgaaccata gggagctgat cgtaat 526

<210> 20
 <211> 516
 <212> DNA
 <213> Homo Sapiens

<400> 20
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 ccatcatccc cgagctggag cgtgagatcc gcatcatcaa cacggagcag tacatgact 120
 cgctgacgtg gcagcaggcg ctcacggggc tgctggagcg catgcagacc tatcaggacg 180
 cggagtcyag gcagggtgctg gctgcctgga tgaaagagcg gcaggagctg aggtgcatca 240
 ccaaggccct gttaaatgcg cagttcggca gcatcttcg caccttccac aacccacct 300
 acttctcaag gcgcctcgtg cgcttctctg acctctacat ggcctccctc agctgcctgc 360
 tcaactaccg cgtggacttc acctctacc cagcccgtag gccgctgcag cagagggcac 420
 ccctctggat ggaccagctc tgcaaccggt gcatgaagac ccccttcctt ggtgacatgg 480
 cccacatccg ctgagggcac ctttattgtc tgggac 516

<210> 21
 <211> 482
 <212> DNA
 <213> Homo Sapiens

<400> 21
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 cagtctaaat tctttcactt acatctttac agaaaactat attttctctc ttccataccc 120
 agaaatctaa tcagaaaact gacttttctc atgttcaact ggacctaggg gaatatgaca 180

8

gaaaagcatc ccataggctt taatatactt tttaaaatat ataaaactga aaattaatag 240
 ccatattaccc tgaaagagtt ctgctgtggac tttgtcactt gcatagtaat agcatgtgcc 300
 tcattgttca gaagattagc tttaggctct attttcaa atcgaaatgg agcataagct 360
 gtaaaactgt agtcttctct gcagaaaata aaggccaaca ataagaaagc ttttgaagga 420
 atcacggaaa acaaatttat aaaagaaata actatatgcg cagtaattct taacacattg 480
 ac 482

<210> 22
 <211> 459
 <212> DNA
 <213> Homo Sapiens

<400> 22
 gcaagtcgcg tgatttctac cacacctgct actgcctgag cggcctgtcc atagcccagc 60
 acttcggcag cggagccatg ttgcatgatg tggctcctggg tgtgcccga aacgctctgc 120
 agcccaactca cccagtgtac aacattggac cagacaaggt gatccaggcc actacatact 180
 ttctacagaa gccagtccea ggttttgagg agcttaagga tgagacatcg gcagagcctg 240
 caaccgacta gaggacctgg gtcccggcag ctctttgctc acccatctcc ccagtcagac 300
 aaggtttata cgtttcaata catactgcat tctgtgctac acaagcctta gcctcagtgg 360
 agctgtgggt ctcttggtac tttcttgtea aaaaaacca atggctctgg gtttgagaa 420
 cacagtggct ggttttaaaa ttctttccac acctgtcaa 459

<210> 23
 <211> 549
 <212> DNA
 <213> Homo Sapiens

<400> 23
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 cctcacaagc agtgacacct cgggtccttt ccgttgctat ggtgaaaatt cctggatgga 120
 atggatcaca tgagggtttc ttgttgcttt tggagggtgt gggggatatt ttgttttgg 180
 ttttctgcag gttccatgaa aacagccctt ttccaagccc attgtttctg tcatggtttc 240
 catctgtcct gagcaagtca ttctttgtt atttagcatt tcgaacatct cggccattca 300
 aagcccccat gttctctgca ctgtttggcc agcataacct ctagcatcga ttcaaagcag 360
 agttttaacc tgacggcatg gaatgtataa atgagggtgg gtccttctgc agatactcta 420
 atcactacat tgctttttct ataaaactac ccataagcct ttaaccttta aagaaaaatg 480
 aaaaagggtta gtgtttgggg gccgggggag gactgaccgc ttcataagcc agtacgtctg 540
 agctgagta 549

<210> 24
 <211> 372
 <212> DNA
 <213> Homo Sapiens

<400> 24
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 cacctactct tatttactgc ccattgattg acttttcttc atattttgca aagagaaatt 120

9

tcacagcaaa aattcatgtt ttgtcagctt tctcatgttg agatctgtta tgtcactgat 180
 gaatttaccc tcaagtttcc ttctctgtga ccactctgct tccttggaca atatcagtaa 240
 tagctttgta agtgatgtgg acgtaattgc ctacagtaat gaaaaattaa tgtactttaa 300
 tttttcattt tcttttagga tatttagacc acccttggtc caccgaaacc agagtgtgtc 360
 agtgtttgtg tg 372

<210> 25
 <211> 475
 <212> DNA
 <213> Homo Sapiens

<400> 25
 cagggatcgg aggacgaccc gagtcccaag agtgggggtt tgcttttttaa aaggagagag 60
 gaggggtgat ggcaggggag tggaggggtg cgggagcagg cctgccggcg cagggagccc 120
 tctgcccttc acactctcct ccaaaagagc ctccatctgt aaggagcag gtctccgca 180
 ggggtttctt tccatgtgtt ttctctctgt tgttaaaaga acttttttaa aaaaacagac 240
 ctctgttttag atttatagca ttgactttta cacacattca cacaagaaaa aaatcctttc 300
 aaaattctta aatcttctgt tctctctttt tccaagggaa gagggcaaaa agtggcctgg 360
 gctctgttgg tgtgcgtgtt ccgtggcgga gagaagaaaa tgggaaagac atctcactgg 420
 tgcttttctc ttttgtttta gtgcccccg ccccatccc tataatatct gtaac 475

<210> 26
 <211> 516
 <212> DNA
 <213> Homo Sapiens

<400> 26
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 ttacactttt atttctgtga tgtgtacaa tcaaaacaca ctactacctc ttaagtccca 120
 gtatacctca tttttcatac tgaaaaaaaa agcttgtggc caatggaaca gtaagaacat 180
 cataaaattt ttatatatat agtttatttt tgtgggagat aaattttata ggactgttct 240
 ttgctgttgt tggctgcagc taaataagac tggacattta acttttctac catttctgca 300
 agttaggtat gtttgccagg agaaaagtat caagacgttt aactgcagtt gactttctcc 360
 ctgttccttt gagtgtcttc taactttatt ctttgttctt tatgtagaat tgctgtctat 420
 gattgtactt tgaatcgctt gacttgttga aaatatttct ctagtgtatt atcactgtct 480
 gttctgcaca ataaacataa cagcctctgt gatccc 516

<210> 27
 <211> 566
 <212> DNA
 <213> Homo Sapiens

<400> 27
 gcgttttccaa cctcggagaa ttccaggcac tccccttccc cctccgctga catacttgta 60
 taagcgggtca tcgttgcgtc atggggcagg cgtggggagc ttctgtcgc cttggctggg 120
 tgtgggcctg gaggaaggtc ctggggcgtg cactgcctg ggcagtgggg aggagagtgg 180

10

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cctgagttac ttcacccccg cgtgctgctg gttaatgtcc cgcgtctctg caccttcggg 240
tgaggagcggg gactgatcta ctttcacatt ctcaagtttt tctcatctgc attagaggtc 300
cccagtaggt tcccagggtc cagcgtgccc ctccctcaga cacacggaca caatcagccg 360
agaagttcct ggtctgaatc acgagaatgt ggaggggtgg ggggtgtcag tggaaaggca 420
taaggctgag ctgagaccag ttgctggtga aactgggcca atctggggag gggaacatcc 480
ttgccaggga gtttctgagg gtctgctttg tttacctttc gtgcggtgga ttctttttaa 540
ctccgtctac ctggcgtttt gttaga 566

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<210> 28
<211> 327
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (199)..(199)
<223> n is a, c, g, or t

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<400> 28
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aagctctggg tgacacgtgg ctccagatca aagcggccaa ggacggagca tccagcagcc 180
ctgagtcgcc ctctcgcan ggcagcccg ccagccctc tgcccacatg gtcagccaca 240
gtcactcccc ctctgtgtgc tctgaaggg agcgctcct ccaacaacac gtggatctgc 300
atggtttgcc tgagctttga acagtca 327

```

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<210> 29
<211> 347
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (156)..(156)
<223> n is a, c, g, or t

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<400> 29
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tccagttagt gacttttgcc ccatagttgg ggtaancact tcctagattg agaaaaagca 180
gctacagtca atcctgctct gtttgctca tttggtgatc agtcagtcac acataagtcc 240
cttgtattct aaatttcatg cacttctccc agatgctata gggttttctc tcactgttgc 300
caatggatgt catccagaca gtgggctcat atcttacggg tttgtgc 347

```

```

<210> 30
<211> 210
<212> DNA
<213> Homo Sapiens

```

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<400> 30
agttgatcag agccttccag agtgtggtat gcttttccact gtgtgatgat ccttagtggc 60

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11

acatgaatga acgtccagat gtttgtgcag tagccacccc ttatctgcag gatacgttcc 120
 aagacccccca gtgaatgcct gaaactgcag atagtactga atcctatata tactgtgttt 180
 tttatgatac atacatgcct atgatgaagt 210

<210> 31
 <211> 511
 <212> DNA
 <213> Homo Sapiens

<400> 31
 aagagaatgt tcctactcac acttcagctg ggtcacatcc atccctccat tcatccttcc 60
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 ctgtgtgccca ggggctggtg ggacagtggg gacatagtct ctgccctcat agagttgatt 180
 gtctagttag gaagacaagc attttttaaaa aataaattta aacttacaaa ctttgtttgt 240
 cacaagtggg gtttattgca ataaccgctt ggtttgcaac ctctttgctc aacagaacat 300
 atgttgcaag accctcccat gggggcactt gagttttggc aaggctgaca gagctctggg 360
 ttgtgcacat ttctttgcat tccagctgtc actctgtgcc tttctacaac tgattgcaac 420
 agactgttga gttatgataa caccagtggg aattgctgga ggaaccagag gcacttccac 480
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<210> 32
 <211> 505
 <212> DNA
 <213> Homo Sapiens

<400> 32
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 aaaaaactgt ttagaattt cttaatttgg atctatttat tagtcagagt ttcagctttc 120
 ttcagctgcc agtgtgttac tcatctttat cctaaaaatc tggaatcaga gatttttgtt 180
 tgttcacata tgattctctt agacactttt atatttgaaa aaattaaaaat ctttctttgg 240
 ggaaaaatcc ttggttatcc tgccataaca gattatgtat taactttagat attcagtggg 300
 tcaatacctg tttagtgtgt tgctaataatt tccagaagga tttcttgtat tggtgaaaga 360
 cggttgggga tgggggggatt tttttgttct tgttgtacct ttgttttgaa actagaaatc 420
 tgtcctgtgg catgcaaaag aaagcaaatt atttttaaaa gaaaaaaacc aaagtacttt 480
 tgggtgcatt attccatctt ctcca 505

<210> 33
 <211> 307
 <212> DNA
 <213> Homo Sapiens

<400> 33
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 acttgggtat ttaaaccatg tcagagctgc ctaccacag gaccttgctg gaggtatac 120
 ttcttctctt gcttgcaca gagcactaca ggatgcattc agtgggcttt tctggcagcc 180
 cagttaacca ttataagat ttggaccttg gagctgaacc agggagctag caaaagtaaa 240

12

gcagacttat aaaattatag ctatgtgcag ctgcacaaca cagtccttcc actagcagct 300
gtgttaa 307

<210> 34
<211> 519
<212> DNA
<213> Homo Sapiens

<220>
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<222> (130)..(130)
<223> n is a, c, g, or t

<220>
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<222> (144)..(144)
<223> n is a, c, g, or t

<220>
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<222> (167)..(167)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (169)..(169)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (268)..(268)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (349)..(349)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (358)..(358)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (478)..(478)
<223> n is a, c, g, or t

<400> 34
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ggaacgcggg cggtgcggac tcagcgggcc ggggtgcaggc gcggagctgg gcctctgcgc 120
ccggcccgan ctccgtctat aaanagagca gccagttgca gggctcnant ctgctttcca 180
actgectgac tgcttggtcg tctcactggt gtgagctoca gcatccctt tgctcgaaat 240
ggaccccaac tgctcttgcg ccaactggntg gctcctgcac gtgcgccggc tcctgcaagt 300
gcaaagagtg caaatgcacc tcctgcaaga agagctgctg ttcctgctnc ccctgggnct 360
gtgccaagtg tgcccagggc tgcgtctgca aaggggcacg ggagaagtgc agctgctgtg 420
cctgatgtgg gaacagctct tctcccagat gtaaatagaa caacctgcac aacctggnat 480
ttttttaaaa atacaacact gagccatttg ctgcatttc 519

<210> 35

13

<211> 460
 <212> DNA
 <213> Homo Sapiens

<400> 35
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 taagggtgaaa aggggtgtgg caaaggagcc tactccaaga atggaggctg taggaatata 120
 acctcccacc ctgcaaaggg aatctcttgc ctgctccatc tcataggcta agtcagctga 180
 atccccgatag tactagggtcc ccttccctcc gcatcccgtc agctggaaaa ggccctgtggc 240
 ccagaggctt ctccaaaggg aggggtgacat gctggctttt gtgccaagc tcaccagccc 300
 tgccgccacct cactgcagta gtgcaccatc tcaactgcagt agcacgccct cctgggcccgt 360
 ctggcctgtg gctaattggag gtgacggcac tcccatgtgc tgactcccc catccctgcc 420
 acgctgtggc cctgcctggc tagtccttgc ctgaataaag 460

<210> 36
 <211> 540
 <212> DNA
 <213> Homo Sapiens

<400> 36
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 atgctgggtg tgatttttta gcccctaaat aaaacactgg actatttccct gtttacttca 120
 ttgattgcaa ctacaaagggt ggactcaaag caaagcacia tcatgccagc caacattcca 180
 gaattctgct gagaactcca agtctgtgag gggagagggt ttacaagcca gacaggcctg 240
 ggggactgca gtccccaagg agaccctgcc acatgctggc cctttgagtg agaattgctgc 300
 atctttctac atatcttcat gagaatactg agaattggat tttccttttc aaaatgcact 360
 ttgctttttt tgtatgtttt gttatgttga gatgtttcta aagaaaagat tttatgtaat 420
 tataagatga agcgtagtga attgtacagc tgttgaata atgacctatt tctatataaa 480
 ataaaattgt atggcttatg tgtaaattat tttgtatctg agataccagt tccttttccc 540

<210> 37
 <211> 367
 <212> DNA
 <213> Homo Sapiens

<400> 37
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 gatgctggcc acaggcgagt gtactcctga ggatctgtgt ttctccctgc aggaaactgt 120
 gtttgcaatg ctggtagaga tcacagagcg agccatggca cattgtggct ccaggaggcc 180
 cctcattgtg ggaggagtgg ggtgtaatgt gaggctacag gagatgatgg caacaatgtg 240
 ccaggaacgt ggagcccggc tttttgctac agatgagaga ttctgtattg acaatggagc 300
 gatgatagcc caggctggct gggagatggt tcgggctgga cacaggacce cactcagtga 360
 ttctggg 367

<210> 38
 <211> 532
 <212> DNA
 <213> Homo Sapiens

14

<400> 38
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 agtccttcac cttgccaggt gccgtttctc ttccgtgaag gccactgcc aggtccccag 120
 tgcgccccct agtggccata gcctgggtaa agttccccag tgctccttg tgcataagacc 180
 ttctttctcc accccttct gccctgggt ccccgccat ccagcggggc tgccagagaa 240
 cccagacct gcccttacag tagtgtagcg cccctccct ctttcggctg gtgtagaata 300
 gccagtagtg tagtgcggtg tgcttttacg tgatggcggg tgggcagcgg gcggcgggct 360
 ccgcgcagcc gtctgtcctt gatctgccg cggcgggccg tgtgtgttt tgtgctgtgt 420
 ccacgcgcta aggcgacccc ctccccgta ctgacttctc ctataagcg ttctcttcgc 480
 atagtcaagt agtcccacc ccacctctt cctgtgtctc acgcaagttt ta 532

<210> 39
 <211> 551
 <212> DNA
 <213> Homo Sapiens

<400> 39
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 ggtgccgttt ctcttccgtg aaggccactg ccaggtccc cagtgcgccc cctagtggcc 120
 atagcctggt taaagttccc cagtgcctcc ttgtgcatag accttcttct cccaccccct 180
 tctgccctg ggtccccggc catccagcgg ggctgccaga gaaccccaga cctgccctta 240
 cagtagtgta gcgccccctc cctctttcgg ctggtgtaga atagccagta gtgtagtgcg 300
 gtgtgctttt acgtgatggc ggggtggcag cggcgggcgg gctccgcgca gccgtctgtc 360
 cttgatctgc ccgcgccggc ccgtgttggtg ttttgtgctg tgtccacgcg ctaaggcgac 420
 cccctcccc gtactgactt ctccataag cgcttctctt cgcatagtca cgtagctccc 480
 accccacct cttctgtgt ctacgcgaag ttttatactc taatatatat atggcttttt 540
 ttcttcgaca a 551

<210> 40
 <211> 538
 <212> DNA
 <213> Homo Sapiens

<400> 40
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 gctcgaagca ggtttaggac caggtccctc tgagaggtca gaggggcctc tgtgggtgct 120
 gggtagtcca gaggtgccac tgggtggaagg gtcagcggag cccagtgcc tccttgtagc 180
 tagac-ttct tctccacccc cttctgccc ctgggtcccc ggccatccag cggggtgcc 240
 agagaacccc agacctgcc ttacagtagt gtacgcgccc ctccctcttt cggctgggtg 300
 agaatagcca gtagttagt gcggtgtgt tttacgtgat ggcgggtggg cagcgggcgg 360
 cgggctccgc gcagccgtct gtccttgatc tgcccgcggc ggcccggtgt gtgttttgtg 420
 ctgtgtccac gcgctaaggc gacccccctc cccgtactga cttctctat aagcgttct 480
 ctctgcatag tcacgtagct cccaccccac cctcttctg tgtctcagc aagtttta 538

15

<210> 41
 <211> 403
 <212> DNA
 <213> Homo Sapiens

<400> 41
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 ccatccggaa cagggccttc tacatggagg agggcgtgcc ctattgagag cgagactatg 120
 agaagatgtt tggcacgaaa tgccatggct gtgacttcaa gatcgacgct ggggaccgct 180
 tcctggaggc cctgggcttc agctggcatg acacctgctt cgtctgtgag atatgtcaga 240
 tcaacctgga aggaaagacc ttctactcca agaaggacag gcctctctgc aagagccatg 300
 ccttctctca tgtgtgagcc ccttctgccc acagctgccg cgggtggcccc tagcctgagg 360
 ggcttgaggt cgtggccctg catttctggg tagggtggc aat 403

<210> 42
 <211> 437
 <212> DNA
 <213> Homo Sapiens

<400> 42
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 gcaggcctgt ggactttctc ctcgagacat taccactatt aaacttctca atgaaactag 120
 agacatgttg gaaagcccag atttttagtac agttttgaat acctgtttaa accgagggtt 180
 tagtagactt ctagacaata tggtgagtt ctttcgacct actgaacagg acctgcaaca 240
 tggtaactct atgaatagtc tttccagtgt cagcctgcct ttagctaaga taattccaat 300
 agtaaacgga cagatccatt cagtttgagc tgaacacct agtcattttg ttcaggatct 360
 gttgacaatg gagcaagtga aagactttgc tgctaagtgt tatgaagctt ttagtacccc 420
 tcagcaactg gagaaat 437

<210> 43
 <211> 520
 <212> DNA
 <213> Homo Sapiens

<400> 43
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 cggattccgg catccacatc atcctccgca ctgagtgagg gtggggagcc caggcctggc 120
 ctggggcag ggcaggggcg ctagggcggc cagctcccc ttgcccgcga gccagtggcc 180
 gaacccccca ctccctgcca ccgtcacaca gtattttattg ttcccacaat ggctgggagg 240
 gggcccttcc agattggggg ccctggggtc cccactccct gtccatcccc agttggggct 300
 gcgaccgcca gattctccct taaggaattg acttcagcag ggggtgggagg ctcccagacc 360
 cagggcagtg tgggtgggagg ggtgttccaa agagaaggcc tggtcagcag agccgccccg 420
 tgtcccccca ggtgctggag gcagactcga gggccgaatt gtttctagtt aggccacgct 480
 cctctgttca gtcgcaaagg tgaacactca tgcggcagcc 520

<210> 44
 <211> 530

16

<212> DNA
 <213> Homo Sapiens

<220>
 <221> misc_feature
 <222> (68)..(68)
 <223> n is a, c, g, or t

<400> 44
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 tgtagacntt tgttgatatt cttcctggaa gaatatcatt cttttcttga agggttgggt 120
 tactagaata ttcaaatca atcatgaagg cagttactat tttagagtcta aagggtttct 180
 aaaaattaac ctacatccc ttctgttagg gtctttcaga atatctttta taaacagaag 240
 catttgaagt cattgctttt gctacatgat ttgtgtgtgt gaaggacata ccacgtttaa 300
 atcattaatt gaaaaacatc atataagccc caactttgtt tggaggaaga gacggagggt 360
 gaggtttttc cttctgtata agcacctact gacaaaatgt agaggccatt caaccgtcaa 420
 acaccatttg gttatatcgc agaggagacg gatgtgtaaa ttactgcatt gctttttttt 480
 tcagtttgta taacctctaa tctccgttg catgatacgc tttgttagaa 530

<210> 45
 <211> 485
 <212> DNA
 <213> Homo Sapiens

<400> 45
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 ggtctggttg atcgatgtca gtcagtcagt agaacctacc cacctcacg gcctggaggt 120
 cttgttccgg gactgcagga atgtctcgca gtttttccag aaaggaggag tcaaggaagc 180
 ccttagtgaa cgagaactct tcaatgctgt ttcaggetta aacatcacag cagataatga 240
 agctgatttt ttagctgaga tagaagcttt ggagaaaatg aatgaagatc acgttcagaa 300
 gaatggaagg aaagctgctt cttttttgaa agatgatgga gaccaccac tactatatga 360
 tgaatagcac taataccac tgcttcagt ttaacacagc agtgattgtc agctgccaat 420
 agcaaatgaa gttatgggtg acttgaaata ccaaacctg aggagtgggc aatggtgctt 480
 ctgtg 485

<210> 46
 <211> 351
 <212> DNA
 <213> Homo Sapiens

<400> 46
 ttgcgccat tggccgtgtt ggtcttgaac tcttgccctc aagcaatccg cctacctcag 60
 cctcccaaag tgctaggatt acaggcataa gccactgagc ccagccctag ttcagtatct 120
 tttatgtaaa ttataaacat ctgcaacatt atgtatcata tgcagatact tattgcattt 180
 cttttattag tgggtgaaagt gttctatgca tttattggct cttgaatttc ctcatctatg 240
 aattgtcatt cacacaccta cttttctgct tcgtttttac atatgtcttt gcctattaaa 300
 gatattatcc ctctgtttta tttttctct cattcttgta ttgcctttta a 351

17

<210> 47
 <211> 521
 <212> DNA
 <213> Homo Sapiens

<400> 47
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 ctctctctcc aacaagcaga accaactctc tcctctggaa gggggcaagc cgctcatgtc 120
 cagctcagaa gaggaattct cacctcccca aagtccagac cagaactcgg tccttctgct 180
 gcagggcaat atggggccacg ccaggagctc aaactattct ctcccgggct taacagcctc 240
 gcagcccagt cacggcctgc agaccaccca gcatcagctc caagactctc tgctcggccc 300
 cctcacctcc agtctgggtg acttggggtc ctaagtgggg agggactggg gcctcgaagg 360
 gattcctgga gcagcaacca ctgcagcgac tagggacact tgtaaataga aatcaggaac 420
 atttttgcag cttgtttctg gagttgtttg cgcataaagg aatggtggac tttcacaaat 480
 atctttttta aaatcaaaac caacagcgat ctcaagctta a 521

<210> 48
 <211> 498
 <212> DNA
 <213> Homo Sapiens

<400> 48
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 ctgtgcccgc aaaccagaca cataggacaa agtttatcta taacctggaa gaccatgagt 120
 ggtgtgaaaa catggagtcg gttttatagt gactaaagga gggctgaact ctgtattagt 180
 aatccaaggg tcattttttt cttaaaaaaa gaaaaaaagg ttccaaaaaa aacccaaact 240
 cagtacacac acacaggcac agatgcacac acacgcagac agacacaccg actttgtcct 300
 ttttctcagc atcagagcca gacaggattc agaataagga gagaatgaca tcgtgcggca 360
 gggctcctgga ggccactcgc gcggctgggc cacagagtct actttgaagg cacctcatgg 420
 ttttcaggat gctgacagct gcaagcaaca ggactgcca aattcaggga acagtgggtg 480
 ccagcttgga ggatggac 498

<210> 49
 <211> 331
 <212> DNA
 <213> Homo Sapiens

<400> 49
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 aaggtaacag gcatctttct tctcagtgcc ttggccctgt tgagtctatc tggtaacact 120
 ggagctgact ccctgggaag agaggccaaa tgttacaatg aacttaatgg atgcaccaag 180
 atatatgacc ctgtctgtgg gactgatgga aatacttate ccaatgaatg cgtgttatgt 240
 tttgaaggtc ggaaacgcca gacttctatc ctcatc aaa aatctggggc ttgctgagaa 300
 ccaaggtttt gaaatcccat caggtcaccg c 331

<210> 50
 <211> 548

<212> DNA
 <213> Homo Sapiens

<400> 50
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 attctgagat gtgcaaagac taccgagtat tgcccaggat aggtctatctt tgtccaaagg 120
 atttaaagcc tgtctgtggt gacgatggcc aaacctacaa caatccttgc atgctctgtc 180
 atgaaaacct gatacgccaa acaatacac acatccgcag tacaggaag tgtgaggaga 240
 gcagcacccc aggaaccacc gcagccagca tgccccgctc tgacgaatga caggaagatt 300
 gttgaaagcc atgagggaaa aaataaaccc cagttctgaa tcacctacct tcacctctg 360
 tatatacaaa gaattcttcg gagcttgtct tatttgctat agaaaacaat acagagcttt 420
 tgggaatgga atcactgatt ttcagtcttt tccatttctt tcctcctaga atctgtgatc 480
 tgaggggtata aagacatttc caccaagttt gagccctcaa aatgtcctga ttacaatgct 540
 gtctgtcc 548

<210> 51
 <211> 526
 <212> DNA
 <213> Homo Sapiens

<400> 51
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 gtcattaaaa atacaccctc tacttgggct ttatactgca taaaaattta ctcatgagcc 120
 ttcctttgag gaaggatgtg gatctccaaa taaagattta gtgtttattt tgagctctgc 180
 atcttaacaa gatgatctga acacctctcc tttgtatcaa taaatagccc tggtattctg 240
 aagtgagagg accaagtata gtaaaatgct gacatctaaa actaaataaa tagaaaacac 300
 caggccagaa ctatagtcac actcacacaa agggagaaat ttaaactcga accaagcaaa 360
 aggttcacg gaaatagcat ggaaaaacaa tgcttccagt ggccacttcc taaggaggaa 420
 caaccccgctc tgatctcaga attggcacca cgtgagcttg ctaagtgata atatctgttt 480
 ctactacgga tttaggcaac aggacctgta cattgtcaca ttgcat 526

<210> 52
 <211> 476
 <212> DNA
 <213> Homo Sapiens

<400> 52
 tgggggactt atttgttggg gatcttaaat aagattcctt ttgatctacc ggaatatata 60
 tgtacagagt acattggatc atgttggaaa gaaggcaagt gaaaaggtca gagatgaagt 120
 agcgaagtta tggaatatcg tggaaaggat actagtgtg aaatggaaag agacaagttt 180
 tagtacccca aaagcaaaac aagcaggaga tgcaagagat gccccaaaag gacaaagcaa 240
 caattttctg ttgccacctt tataccggaa gactctgttg tagaagaaaa gaaggctttg 300
 gtgcacctta tgtgggagga ggaggggagc ggcattgctga tgctgagcgt acaggcagac 360
 aagagcgtag cctgctgttg cctccatcac tatgaaatga cttattttac ctgaaggacc 420
 catggtttat gtctctctaa ttcctttcac tctccctaag ccctctgaga gagatg 476

19

<210> 53
 <211> 501
 <212> DNA
 <213> Homo Sapiens

<400> 53
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 ataagaaccg tgtacggagt cgtatctcca acctgaagga tgccaagaac cctgacctgc 120
 ggcggaatgt gctgtgtggg gccataacac cccagcagat cgctgtgatg acctcagagg 180
 agatggccag tgatgagctg aaggagatcc gtaaggccat gaccaaggag gccatccgag 240
 agcaccagat ggcccgcact ggccgcacgc agacagacct gttcacctgc ggcaagtgc 300
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 cctttgttgt ctgcaacgag tgtggaaacc gctggaagtt ctgctgacct ctctgttaga 420
 tgtgtctgcag ccttggggccc tccccggccc acgtcctccg ttgacacagc ttctctggag 480
 accctagaag gggcatgtc c 501

<210> 54
 <211> 453
 <212> DNA
 <213> Homo Sapiens

<400> 54
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 tgggcaagyt gatgaggatg ctgggcccaga accccacccc tgaggagctg caggagatga 180
 tcgatgaggt ggacgaggac ggcagcggca cgggtggactt tgatgagttc ctggtcatga 240
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 gcatgtttga caaaaatgct gatggctaca tcgacctgga tgagctgaag ataatgctgc 360
 aggctacagg cgagaccatc acggaggacg acatcgagga gctcatgaag gacggagaca 420
 agaacaacga cggccgcac cactatgatg agt 453

<210> 55
 <211> 498
 <212> DNA
 <213> Homo Sapiens

<220>
 <221> misc_feature
 <222> (142)..(142)
 <223> n is a, c, g, or t

<400> 55
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 caagttcccc ggagagtgg gntccctgt gggccctca ggcctatgtc tgtgaggaag 180
 gggccctgcc actctcccca agagggcctc catgtttcga ggtgcctcaa catggagcct 240
 tgcttggcct gggctagggg cactgtctga actcctgact gtcaggataa actccgtggg 300
 ggtacaggag cccagacaaa gccaggcct gtcaagagac gcagagggcc cctgccaggg 360

20

ttggccccag ggaccctggg acgaggctgc agaagctctc cctccctact cctggggagc 420
 cacgtgctgg ccatgtggcc agggacggca tgagcaggag gcggggacgt gggggccttc 480
 tggtttggtg tcaacagc 498

<210> 56
 <211> 544
 <212> DNA
 <213> Homo Sapiens

<400> 56
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 ggagaagcaa tatgagtgac tgaagagtgg ggctttttgc ttttgcctgg atataggggt 120
 gctcttctac tgtaattggg tgtggaaaaa ctctggcttt atggtattcc attaggttct 180
 tttcatttaa agtagtctta aaatcaaagt atccaatatt ttaaagccac aaagtagatt 240
 acataattag cagagatttt agtcagtaaa atgttagaaa tcaaaactata agaaaattca 300
 agtcctttat tttgtgtctt gggatatatgt cattatttta aattccacac tcccttattt 360
 aatcactttg gtaagtgcct ttgatgtttt gaaatgtata gtgggagatg agcaaagtga 420
 aatgtcatgt gccctgttcc ctagcttctc aattcctcat aaccattttt accagtgttg 480
 caaagtttag acctttgtgt taatatcaga agtgtatttg tagccctcc atagtgaaca 540
 atga 544

<210> 57
 <211> 535
 <212> DNA
 <213> Homo Sapiens

<400> 57
 gccgaagaag cctgtctgtg ggggtgtgtc cagcgctctg gcacctggcg tccgagccgt 60
 ggagctcgag cggcagatcg agagcacaga gacttcttgc catggctgcc gtaagaattt 120
 cttctgttcc aagatccggt cccacgtggc tacttgttcc aaataccaga attacatcat 180
 ggaagggtgtg aaggccacca ttaaggatgc atctcttcag ccaaggaatg tcccaaaccg 240
 ttacaccttt ccttgtcctt actgtcctga gaagaacttt gatcaggaag gacttgtgga 300
 acactgcaaa ttattccata gcacggatac caaatctgtg gtttgtccga tatgtgcctc 360
 gatgccctgg ggagacccca actaccgag cgccaacttc agagagcaca tccagcgccg 420
 gcaccgggtt tcttatgaca cttttgtgga ttatgatgtt gatgaagagg acatgatgaa 480
 tcagggtgtg cagcgctcca tcatcgacca gtgagcagag tccgtgcttg ctatc 535

<210> 58
 <211> 479
 <212> DNA
 <213> Homo Sapiens

<400> 58
 gctgaaagaa gcccacatag aactgcttag ggacagcacc actgactcca aagaaaatcc 60
 cagcagaaag agaaatggaa tgtgcacgga tacacattca ctgctcagta agaggctcaa 120
 gacatgactg atttgcattt taaagcaaga tgcgatgtcc agagttacag agaattagta 180

21

gatgtgtctc atcggttaaat agctctatta tacctctaaa ggtggaattg tcagttttaga	240
ttcataaatg aaaaggtaaa tgagtaatca gaataaacca agtgataatc aaacctgtc	300
aagattatta gttcagactc tagcctgtta attttcttag ttgatttctg aagctacctg	360
atattattcta ttaaattgta agcttgcaaa ctcaaaataa attggcagat ttacctctca	420
tgttttaatg tgtcaaatta gagagcaaag tataacaggt gccttcactt ttgagactt	479

<210> 59
 <211> 518
 <212> DNA
 <213> Homo Sapiens

<400> 59 gtgccatagt gcaggcttgg ggagctttaa gcctcagtta tataaccac gaaaaacaga	60
gcctcctaga tgtaacattc ctgatcaagg tacaattctt taaaattcac taatgattga	120
ggtccatatt tagtggtact ctgaaattgg tcactttcct attacacgga gtgtgctaaa	180
actaaaaagc attttgaaac atacagaatg ttctattgtc attgggaaat ttttctttct	240
aacctcagtgg aggttagaaa gaagttatat tctggtagca aattaacttt acatcctttt	300
tcctacttgt tatggttggt tggaccgata agtgtgttta atcctgaggc aaagtagtga	360
atatgtttta tatgttatga agaaaagaat tggtgtaagt ttttgattct actcttatat	420
gctggactgc attcacacat ggcatgaaat aagtcagggt ctttacaat ggtattttga	480
tagatactgg attgtgtttg tgccatattt gtgccatt	518

<210> 60
 <211> 489
 <212> DNA
 <213> Homo Sapiens

<400> 60 gggatgcatt tgtggccatt gttcaaagt tcaagaacaa gcctctcttc ttgccgaca	60
aactttacaa atccatgaag ggtgctggca cagatgagaa gactctgacc aggatcatgg	120
tatcccgcag tgagattgac ctgctcaaca tccggaggga attcattgag aaatatgaca	180
agtctctcca ccaagccatt gagggtgaca cctccggaga cttcctgaag gccttgctgg	240
ctctctgtgg tggtagaggac tagggccaca gctttggcgg gcacttctgc caagaaatgg	300
ttatcagcac cagccgccat ggccaagcct gattgttcca gctccagaga ctaagggaagg	360
ggcaggggtg gggggagggg ttgggttggg ctcttatctt catggagctt aggaaacgct	420
cccactcca cgggccatcg agggccagca cggctgagcg gtgaaaaacc gtagccatag	480
atcctgtcc	489

<210> 61
 <211> 472
 <212> DNA
 <213> Homo Sapiens

<400> 61 atttcaaat ttctgcatc acggagaatg caaatatata gagcacctgg aagcagtaac	60
atgcaaatgt cagcaagaat atttcggtga acggtgtggg gaaaagtcca tgaaaactca	120
cagcatgatt gacagtagtt tatcaaaaat tgcattagca gccatagctg cctttatgtc	180

22

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tgctgtgac ctcacagctg ttgctgttat tacagtccag cttagaagac aatacgtcag      240
gaaatatgaa ggagaagctg aggaacgaaa gaaacttcga caagagaatg gaaatgtaca      300
tgctatagca taactgaaga taaaattaca ggatatcaca ttggagtcac tgccaagtca      360
tagccataaa tgatgagtcg gtctctcttc cagtggatca taagacaatg gacccttttt      420
gttatgatgg ttttaaactt tcaattgtca ctttttatgc tattttctgta ta              472

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<210> 62
<211> 523
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (41)..(41)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (440)..(440)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (442)..(442)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (485)..(486)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (488)..(491)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (493)..(498)
<223> n is a, c, g, or t

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<400> 62
gacaacagcc ctggagggga acagagtgag agagatgttt ngctctggta cagcctgtgt      60
tgtttgccca gtttctgata tactgtacaa aggcgagaca atacacattc caactatgga      120
gaatggtcct aagctggcaa gccgcattct gagcaaatta actgatatcc agtatggaag      180
agaagagagc gactggacaa ttgtgctatc ctgaatggaa aatagaggat acaatggaaa      240
atagaggata ccaactgtat gctactggga cagactgttg catttgaatt gtgatagatt      300
tctttggcta cctgtgcata atgtagtttg tagtatcaat gtgttacaag agtgattggt      360
tcttcatgcc agagaaaatg aattgcaatc atcaaatggt gtttcataac ttggtagtag      420
taacttacct taccttaccn anaaaaatat taatgtaagc catataacat gggattttcc      480
tcaannannn nannnnnncc ttttgtactt cactcagata cta                          523

```

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<210> 63
<211> 373
<212> DNA
<213> Homo Sapiens

```


23

<400> 63
 gggcagatct tggactcatg aggaggggcc cccctgccca gaggggtcaa cccttctgga 60
 aactgtgaag atctgacttc gccccccccc cccccatct tcgggaccag gatttgaca 120
 gaagcacatg cacctaccca tacacccct cttctgagcg tccctgttcc cccatctcgc 180
 tccctcccag gactctgacc ccagcattct caggcaccag tccctgtccg gaatgccacc 240
 cacatcttcc atttccatgt cccctcccag agctgggtgga cccaggaac agccactccc 300
 ctccactctc taccagataa ctgaggaggg gagaggtggg ccgtaacggg cacggatcac 360
 gatgtaaatt att 373

<210> 64
 <211> 535
 <212> DNA
 <213> Homo Sapiens

<400> 64
 agcttcagga cgcgtctgca gaggtggagc gactgagaag agaaaaccag gtcttaagcg 60
 tgagaatcgc ggacaagaag tactacccca gctcccagga ctccagctcc gctgcggcgc 120
 cccagctgct gattgtgctg ctgggcctca gcgtctgct gcagtgaat cccaggaagc 180
 tggcacatct tggaaggctc gtctgtctcg gcttttcgct tgaacattcc cttgatctca 240
 tcagttctga gcggtcatg gggcaacacg gttagcgggg agagcacggg gtagccggag 300
 aagggcctct ggagcaggctc tggaggggcc atggggcagt cctgggtgtg gggacacagt 360
 cgggttgacc cagggtgtc tccctccaga gcctccctcc ggacaatgag tccccctct 420
 tgtctccac cctgagattg ggcattgggt gcggtgtggg gggcatgtgc tgctgttgt 480
 tatgggtttt ttttcggggg ggggttgctt tttctgggg tctttgagct ccaa 535

<210> 65
 <211> 452
 <212> DNA
 <213> Homo Sapiens

<400> 65
 catgtctggac cagatcaact cctgtctgga ccacctggag gagaagaatg accacctcca 60
 cgccccctc caggagctgc tggagtccaa ccggcagaca cgctggagt tccagcagca 120
 gctcggggag gccccagtg atgccagccc ctaggctcca agagccccca accgggaccc 180
 aacctgcct ccttgggcta ggctctggcc tgggcaactc cccctggct tagacacctt 240
 ctcaagggct ggccttcagg gaccttggt ggtctgcct gcctgggcca ccttctctgc 300
 ctgggcctcc ccttggcta cctgggccag cccccaccac ctggcatgcc ctctggggc 360
 caagagtggg cctgcaaccc acccacttgc ctgccaccc aactcctggg cgctccccac 420
 tctgccagg ccttgagtgt ccacattaaa tg 452

<210> 66
 <211> 323
 <212> DNA
 <213> Homo Sapiens

<400> 66
 cacttaccag tgagcatata tattttaaaa tactttcttt ggatattgta attcttaact 60

24

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ggttgtaaata tagaaaagct gggattacat atggtgtgcg gttacagtct aaattttttc 120
atcctcctat gcatcataag catgtttgta atattttcaa aaatagttct actgatgcta 180
caggaatttc aagcctgtgg tgaatgtag tatttaccat agggagtga gtggagttag 240
ggtttcattc aatagagtat tgctgattat acttgagtgg aatcctttcc tcacgiactc 300
ccacagacgt ctgggcctgg aaa 323

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<210> 67
<211> 560
<212> DNA
<213> Homo Sapiens

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<400> 67
ggcggaggag aacaaacaga tcatccgcaa acacgcgcag accttcgttg ccctctgtgc 60
cacagatgtg aagttcattt ccaatccgcc ctccatggtg gcagcgggga gcgtggtggc 120
cgcagtgcaa ggcctgaacc tgaggagccc caacaacttc ctgtcctact accgcctcac 180
acgcttcctc tccagagtga tcaagtgtga cccggactgc ctccgggcct gccaggagca 240
gatcgaagcc ctgctggagt caagcctgcg ccaggcccag cagaacatgg accccaaggc 300
cgccgaggag gaggaagagg aggaggagga ggtggacctg gcttgacac ccaccgacgt 360
gcgggacgtg gacatctgag ggcgccaggc aggggggggc caccgccacc cgcagcgagg 420
gcggagccgg cccaggtgct tccccgaca gtccctcctc tccggagcat tttgatacca 480
gaagggaaag cttcattctc cttgtgtgtg gttgtttttt cctttgtctt tttccccctc 540
catctctgac ttaagcaaaa 560

```

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<210> 68
<211> 471
<212> DNA
<213> Homo Sapiens

```

```

<400> 68
gttttgggta tgtttaatct gttatgtact agtgttctgt ttgttattgt tttgttaatt 60
acaccataat gctaatttaa agagactcca aatctcaatg aagccagctc acagtgtgtg 120
gtgccccggt catctagcaa gctgccgaac caaaagaatt tgcacccgcg tcggggccca 180
cgtggttggg gccctgccct ggcagggtca tctgtgtctc ggaggccatc tcgggcacag 240
gccacccccg cccacccct ccagaacacg gctcacgctt acctcaacca tcttggtgct 300
ggcgtctgtc tgaaccacgc gggggccttg agggacgctt tgtctgtcgt gatggggcaa 360
gggcacaagt cctggatggt gtgtgtatcg agaggccaaa ggctggtggc aagtgcacgg 420
ggcacagcgg agtctgtcct gtgacgcgca agtctgaggg tctgggcggc g 471

```

```

<210> 69
<211> 518
<212> DNA
<213> Homo Sapiens

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<400> 69
aattcctgcc attctgggga ttcttgagg aattcttgct ttgctaattc tgattctgct 60
gctcttgctg tttcttcgga ggagagcggg ggtcaaagag cccttactgc cccagagga 120

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25

tgacacccgg gacaacgttt attactatga tgaagaagga ggcggagaag aggaccagga 180
 ctttgacttg agccagctgc acaggggcct ggacgctcgg cctgaagtga ctcgtaacga 240
 cgttgacca accctcatga gtgtcccccg gtatcttccc cgccctgcca atcccgatga 300
 aattggaaat ttattgatg aaaatctgaa agcggtgat actgaccca cagccccgcc 360
 ttatgattct ctgctcgtgt ttgactatga aggaagcggg tccgaagctg ctagtctgag 420
 ctccctgaac tcctcagagt cagacaaaga ccaggactat gactacttga acgaatgggg 480
 caatccgttc aagaagctgg ctgacatgta cggaggcg 518

<210> 70
 <211> 182
 <212> DNA
 <213> Homo Sapiens

<400> 70
 cttttcactg tggttgagtt ttctggagtg agcactcacg ccctaagcgc acattcatgt 60
 gggcatttct tgogagcctc gcagcctccg gaagctgtcg acttcatgac aagcattttg 120
 tgaactaggg aagctcaggg gggttactgg cttctcttga gtcacactgc tagcaaatgg 180
 ca 182

<210> 71
 <211> 538
 <212> DNA
 <213> Homo Sapiens

<400> 71
 tgaggagcca gcgtctaggg cagcagccgc ttctagaag accaggatcat gatgatgggc 60
 agcgcccagag tggcggagct gctgctgctc cagggcggg agcccaactg cgccgacccc 120
 gccactctca cctgacccgt gcacgacgct gcccgggagg gcttccctgga cagctgggtg 180
 gtgctgcacc gggccggggc gcggctggac gtgcgcgatg cctggggccg tctgcccgtg 240
 gacctggctg aggagctggg ccacgcgatg gtgcacggg acctgcgcgc ggctgcgggg 300
 ggcaccagag gcagtaacca tgcccgcata gatgccacgg aaggctccctc agacatcccc 360
 gattgaaaga accagagagg ctctgagaaa cctcgggaaa cttagatcat cagtcaccga 420
 aggtcctaca gggccacaac tgccccgcc acaaccacc ccgcttctgt agttttcatt 480
 tagaaaatag agctttttaa aatgtcctgc cttttaacgt agatatatgc cttcccc 538

<210> 72
 <211> 513
 <212> DNA
 <213> Homo Sapiens

<400> 72
 atattagtta ccctgggtgtg ctgtattctc taaaaccttt aaatgtttgc atgcagccat 60
 tcgtcaaatg tcaaattatc tctctttggc tggaatgaca aaaactcaaa taaatgtatg 120
 attaggagga catcataacc tatgaatgat ggaagtccaa aatgatggta actgacagta 180
 gtgttaatgc cttatgttta gtcaaaactct catcttaggtg acagcctggg gactccagaa 240
 tggagccagt catgctaaat gccatatact cacactgaaa catgaggaag caggtagatc 300
 ccagaacaga caaaactttc ctaaaaacat gagagtccag gctgtctgag tcagcacagt 360

26

aagaaagtcc tttctgcttt aactccttaga aaaaagtaat atgaagtatt ctgaaattaa 420
 ccaatcagtt tatttaaate aatttatatta tattcttctg ttcttggtt cccattttac 480
 aaaaccact gttctactgt tgtattgccc agt 513

<210> 73
 <211> 530
 <212> DNA
 <213> Homo Sapiens

<400> 73
 ggatttgtgt tcttacagta cttgaaaata ttaaggaag agatgaagct ctgcagtttt 60
 ttctatgttg gatgattact tttttaagga ggattaattc tgaggtagta tagtaactaa 120
 aggggaatat atgaattggt taacaaatta gaatttggtt acaactactt gaatttttaa 180
 attatgtcaa aacttacatt acttgccaag cagtatgatg ttataggaaa cataaataag 240
 attacagagg tatcaatttg gttaaaattc accattttat aagactaagc aataatctta 300
 acaacctctt tctgaatat ttaaatgtgt ttgtatggtg ttatgactaa ttgttactga 360
 tttagagact aagccctctt aaaaccttta gttaaatata aaaagaaatt atatatatct 420
 tgcctccctg atggaaaact atataaaatt gtagacttaa aaggtttgtg gaaatacatt 480
 aggatatcag aaaactaaat atatggagtt gctttatgac tattacatgt 530

<210> 74
 <211> 406
 <212> DNA
 <213> Homo Sapiens

<400> 74
 ggctgcctgc ggatgaagga ccagtgtgac aagtgcggg agatcttgtc tgtggactgt 60
 tccaccaaca accctccca ggctaagctg cggcgggagc tgcacgaatc cctccaggtc 120
 gctgagaggt tgaccaggaa atataacgag ctgctaaagt cctaccagtg gaagatgtc 180
 aacacctctt ccttgctgga gcagctgaac gagcagttta actgggtgtc cgggctggca 240
 aacctcacgc aaggcgaaga ccagtactat ctgcgggtca ccacgggtgc tccccacact 300
 tctgactcgg acgttccttc cgggtgcact gaggtggtcg tgaagctctt tgactctgat 360
 cccatcactg tgacggtccc tgtagaagtc tccaggaaga acccta 406

<210> 75
 <211> 286
 <212> DNA
 <213> Homo Sapiens

<400> 75
 agcagctgaa cgagcagttt aactgggtgt cccggctggc aaacctcacg caaggcgaag 60
 accagtacta tctgcgggtc accacggtgg cttccacac ttctgactcg gacgttcctt 120
 cgggtgtcac tgagggtgtc gtgaagctct ttgactctga tccatcact gtgacggtcc 180
 ctgtagaagt ctccaggaag aaccctaaat ttatggagac cgtggcggag aaagcgtgc 240
 aggaataccg caaaaagcac cgggaggagt gagatgtgga tgttgc 286

<210> 76

<211> 436
 <212> DNA
 <213> Homo Sapiens

<400> 76
 gaaagactgt gctgtccttt aacatagggt tttaaagact aggatattga atgtgaaaca 60
 tccgttttca ttgttcactt ctaaaccaaa aattatgtgt tgccaaaacc aaaccagggt 120
 tcatgaatat ggtgtctatt atagtgaaac atgtactttg agcttattgt ttttattctg 180
 tattaaatat tttcagggtt ttaaactact atcacaaact gaatgacttg acttcaaaag 240
 caacaacctt aaaggccgtc atttcattag tattcctcat tctgcatcct ggcttgaaaa 300
 acagctctgt tgaatcacag tatcagtatt ttcacacgta agcacattcg ggccatttcc 360
 gtggtttctc atgagctgtg ttcacagacc tcagcagggc atcgcatgga ccgcaggagg 420
 gcagattcgg accact 436

<210> 77
 <211> 429
 <212> DNA
 <213> Homo Sapiens

<400> 77
 tcggctactc ttttgtgatg cacaccagcg ctggtgcaga aggctctggc caagccctgg 60
 cgtcccccg ctcctgcctg gaggagtta gaagtgcgcc attcatcgag tgtcacggcc 120
 gtgggacctg caattactac gcaaacgctt acagcttttg gctcgccacc atagagagga 180
 gcgagatgtt caagaagcct acgccgtcca cctgaaggc aggggagctg cgcacgcacg 240
 tcagccgctg ccaagtctgt atgagaagaa cataatgaag cctgactcag ctaatgtcac 300
 aacatggtgc tactttctct tctttttgtt aacagcaacg aaccctagaa atatatcctg 360
 tgtacctcac tgtccaatat gaaaaccgta aagtgcctta taggaatttg cgtaactaac 420
 acaccctgc 429

<210> 78
 <211> 195
 <212> DNA
 <213> Homo Sapiens

<400> 78
 tccccctgta gactagtgcc gtgggagtac ctgctgcccc gctgctgtgg cccctccgt 60
 gatccatcca tctccaggga gcaagacaga gacgcaggat ggaaagcgga gttcctaaca 120
 ggatgaaagt tcccccatca gtccccccag tacctccaag caagtagctt tccacatttg 180
 tcacagaaat cagag 195

<210> 79
 <211> 301
 <212> DNA
 <213> Homo Sapiens

<400> 79
 tgggtgtggg agccctttgg agaacgccag tctccaggtc cccctgcac tctcgagttt 60
 gcaatgtcac aacctctctg atcttgtgct cagcatgatt ctttaataga agttttatct 120
 ttcgtgcact ctgctaatca tgtgggtgag ccagtggaac agcgggagcc tgtgctggtt 180

28

tgcagattgc ctctaatga cgcggctcaa aaggaaacca agtggtcagg agttgtttct 240
gacccactga tctctactac cacaaggaaa atagtttagg agaaaccagc ttttactgtt 300
t 301

<210> 80
<211> 459
<212> DNA
<213> Homo Sapiens

<220>
<221> misc_feature
<222> (164)..(164)
<223> n is a, c, g, or t

<400> 80
ggaaacgttc ccagttcatt ttcagtcctg ttgtgagcac agttctgaag ggtttattat 60
tgtcaaaaata agttttgttt tgttttgttt atgttgggtt tttaatgttg tctcttgacc 120
cttaatgctc aggttcttgt gggagttaat cagccacatc caangttacc ttgaggggga 180
agaagagggt gatgctcaga agctaaacaa gacaggggcc acatgaccct ctattgatta 240
gccccaaagta gaaagtcctg tggttttatg tttaatggta atagttgatc atatatggca 300
taattttcta tcagcttcct actcagtcac tataaacaca gacttgaaat agtactttaa 360
atgtccaaat acctaaatgt gctaaactgg aggtaactat ttctaggtag ttgaattttt 420
gaaagtcatg atcagccaca caactgtttt gtacatact 459

<210> 81
<211> 394
<212> DNA
<213> Homo Sapiens

<400> 81
aatccttatt gttcagagtt gtttgggggt tctgtttcag agcataaaac ctaaaggtta 60
tagtagaaca aggcaccttc ttaaaagaaa tcttgcttca gaccatcagt tacagagaat 120
ttcctaaagt aaaattgaag caactacaac ttctccttag acactttgga atctaaccac 180
ttaaggacct ttttaaagag atagcttctc ttctttctga agatcaattt ctccaaggc 240
caagattgtc cttttctccc atttcttgc agctattgca aatgagggaa gaacattatt 300
catctctcct cccctttttt ttctgattct ttttccagtc agttttgtc ctgggttcaa 360
gtagtattac caccctttca caagcaacag actc 394

<210> 82
<211> 514
<212> DNA
<213> Homo Sapiens

<220>
<221> misc_feature
<222> (89)..(89)
<223> n is a, c, g, or t

<400> 82
gtcactaca ctattcattg cacacaaatg aatttttcac tttttaagat gcattcttgg 60
tgctcaaacc agatcgaagt ttgtctctna aagctattgt ctgcacaggc tgctgcatgc 120

29

tctgttggtta aatggatgga caggctattc taaattttgg ttgatacttt tgctactatg 180
ggcaattaac ttgaaaaaaa taatcgatcc caactctgtg ctctgatgta cctcttctgc 240
cccttttatg acacctttga ccaaatgcct tctatgggtc acagtgcagg cacaaaacta 300
cctctgatac agaaggttct ttacaagctt attttacata ccgtgaatcc ctcacctaaa 360
gggagagggtg aaagcaaaga ctgctttgaa tgggtattga gggagattgt gtccatacca 420
agccaccctg aagaagtatt tcaactgcag tagaactgtg gatttgtgct gtcatttcac 480
cttggataaa acacctatct ctaagcagga ccaa 514

<210> 83
<211> 299
<212> DNA
<213> Homo Sapiens

<400> 83
caccaaatta cctaggctga ggtagagag attggccagc aaaaactgtg ggaagatgaa 60
ctttgtcatt atgatttcat tatcacatga ttatagaagg ctgtcttagt gcaaaaaaca 120
tacttacatt tcagacatat ccaaagggaa tactcacatt ttgttaagaa gttgaactat 180
gactggagta aaccatgtat tcccttatct tttacttttt ttctgtgaca tttatgtctc 240
atgtaatttg cattactctg gtggattgtt ctagtactgt attgggcttc ttcgttaat 299

<210> 84
<211> 219
<212> DNA
<213> Homo Sapiens

<400> 84
ttatcgccct gagaagatct accccagggg gaatctgaga catcttgcct acttttcttt 60
attagctttc tctcatcca tttcttttat acctttcctt tttggggagt tgttatgcca 120
tgatttttgg tatttatgta aaaggattat tactaattct atttctctat gtttattcta 180
gttaaggaaa tgttgagggc aagccaccaa attacctag 219

<210> 85
<211> 518
<212> DNA
<213> Homo Sapiens

<220>
<221> misc_feature
<222> (61)..(65)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (71)..(71)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (73)..(73)
<223> n is a, c, g, or t

<220>
<221> misc_feature
<222> (112)..(112)

<223> n is a, c, g, or t

<220>

<221> misc_feature

<222> (163)..(163)

<223> n is a, c, g, or t

<220>

<221> misc_feature

<222> (295)..(295)

<223> n is a, c, g, or t

<400> 85

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aaggactggt atctttctgt gagcaataag gactggataa agactgcata tccttggtgc      60
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ttctggtctt ggatgaataa gttgcacaga gtattgcacc aanaatacac aatggagggt      180
gaaaagttca acatatttta agtcaattaa tcaaattgca ttgattcttg atgctttctt      240
agaggcctac atgatttctt agattgctct gataaactat cataaggggt ccacntcccc      300
tcatttagct cccccagga tttcttttcc cccatgtcat acaccagtc ctaaatcaac      360
ccccagggt atccttccat cccttctgca gaggaactt ttgtcagact ctgcaacaaa      420
ctcctagctc tatccagagt gtccctctgt gctaagattg gtatctttct cctcaaaagc      480
ctggatggtg aatgggggtg cattagtcag aattctcc      518

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<210> 86

<211> 458

<212> DNA

<213> Homo Sapiens

<400> 86

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taaaaaacctg tatctgaccc actttgtaat ttttgtcca atatccattc tgtagacttt      60
tgaaaaaaa gtttttaatt tgatgcccaa tatattctga ccgttaaaaa attcttggtc      120
atatgggaga agggggagta atgacttgta caaacagtat ttctggtgta tattttaatg      180
tttttaaaaa gagtaatttc atttaaatat ctgttattca aatttgatga tgttaaagt      240
aatataatgt attttctttt tattttgac tctgtaattg cactttttaa gtttgaagag      300
ccattttggt aaacggtttt tattaaagat gctatggaac ataaagtgt attgcatgca      360
atttaaagta acttatttga ctatgaatat tatcggatta ctgaattgta tcaatttgtt      420
tgtgttcaat atcagctttg ataatttgtt accttaag      458

```

<210> 87

<211> 336

<212> DNA

<213> Homo Sapiens

<400> 87

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gggatcctat ttagctctta gtaccactaa tcaaaagttc ggcatgtagc tcatgatcta      60
tgctgtttct atgtcgtgga agcaccggat ggggtagtg agcaaatctg ccctgctcag      120
cagtcaccat agcagctgac tgaaaatcag cactgcctga gtagttttga tcagtttaac      180
ttgaatcact aactgactga aaattgaatg ggcaaataag tgcttttgct tccagagtat      240
gcgggagacc ctccacctc aagatggata tttcttcccc aaggatttca agatgaattg      300
aaatttttaa tcaagatagt gtgctttatt ctgttg      336

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31

<210> 88
 <211> 521
 <212> DNA
 <213> Homo Sapiens

<400> 88
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 ataaagtcag aaaataaagt taacataact ttcactaaca cacacatatg tagatttcac 120
 aaaatccacc tataattggt caaagtgggt gagaatatat tttttagtaa ttgcatgcaa 180
 aatttttcta gcttccatcc tttctccctc gtttcttctt tttttggggg agctggtaac 240
 tgatgaaatc ttttccacc ttttctctc aggaaatata agtggttttg tttggttaac 300
 gtgatacatt ctgtatgaat gaaacattgg agggaaacat ctactgaatt tctgtaattt 360
 aaaatatttt gctgctagtt aactatgaac agatagaaga atcttacaga tgctgctata 420
 aataagtaga aaatataaat ttcataccta aaatatgcta ttttaaaatc tatttcctat 480
 attgtatttc taatcagatg tattactctt attatttcta t 521

<210> 89
 <211> 503
 <212> DNA
 <213> Homo Sapiens

<400> 89
 gtggctatcc actgttagtt cagaagctgg gcttggacta ctcttatgat ttagctccac 60
 gagccaaaat tttccggcgt gaccaaggga aagtgactga tacggcatcc atgaaatata 120
 tcatgcgata caacaattat aagaaggatc cttacagtag aggtgacccc tgtaatacca 180
 tctgctgccg tgaggacctg aactcaccta acccaagtcc tggaggttgt tatgacacaa 240
 aggtggcaga tatctaccta gcatctcagt acacatccta tgccataagt ggtccacag 300
 tacaagggtg cctccctgtt tttcgtggg accgtttcaa caaaactcta catcagggca 360
 tgccagaggt ctacaacttt gatatttatta ccatgaaacc aattttgaaa cttgatataa 420
 aatgaaggag ggagatgacg gactagaaga ctgtaaataa gataccaaag gcactatttt 480
 agctatgttt ttcccatcag aat 503

<210> 90
 <211> 275
 <212> DNA
 <213> Homo Sapiens

<400> 90
 ccccatcacg gagggctccag actgtccact cgggggtgga gtgagactga ctgcaagccc 60
 caccctcctt gagactggag ctgagcgtct gcatacgaga gacttggttg aaacttggtt 120
 ggtccttgtc tgcaccctcg acaagaccac actttgggac ttgggagctg gggctgaagt 180
 tgctctgtac ccatgaactc ccagtttgcg aattaataag agacaatcta ttttggtact 240
 tgcacttggt attcgaacca ctgagagcga gatgg 275

<210> 91
 <211> 405
 <212> DNA

<213> Homo Sapiens

<400> 91

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tcattctgatg tttctatagt cactttgcc gctcaaaaga aaacaatacc ctatgtagtt    60
gtggaagttt atgctaatat tgtgtaactg atattaaacc taaatgttct gcctaccctg    120
ttggtataaa gatattttga gcagactgta aacaagaaaa aaaaaatcat gcattcttag    180
caaaattgcc tagtatgtta atttgctcaa aatacaatgt ttgattttat gcactttgtc    240
gctattaaca tccttttttt catgtagatt tcaataattg agtaatttta gaagcattat    300
tttaggaata tatagttgtc acagtaaata tcttgttttt tctatgtaca ttgtacaaat    360
ttttcattcc ttttgctctt tgtggttga tctaacta actgt                        405
```

<210> 92

<211> 375

<212> DNA

<213> Homo Sapiens

<400> 92

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aagctatgtg tatcttctgt gtaaagcagt ggcttactg gaaaaatggg gtggctagca    60
tttccctttg agtcatgatg acagatgggt tgaaaaccat ctaagtttgc ttttgaccat    120
cacctcccag tagcaatttg ctttcataat ccatttagca atccaggcct ctgttgaaaa    180
gataatatga gggagaaggg aacacatttc cttctgaact tacttcctta agtcactttc    240
cttatgtatc atctaataca atgatgggtg agtgaaaata cagaaggggt gtttgagtat    300
tcagatttca taaaacactt ccttggaata tagctgcatt aacttggaag gaagcctggt    360
gggccagaag acaga                                                    375
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<210> 93

<211> 533

<212> DNA

<213> Homo Sapiens

<400> 93

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gctggtgtgt gtgtcaaacc ctcaactcacc cagcactca cacacagcat tctgttctcc    60
atgcaaagtt aagatcgaat ccattcgctt gtaggggaaa aaaaggaaaa aaattaacca    120
gagagggtct gtaatctcgc agagcacagg cagaatcggt ccttccttgc tgcatttcct    180
ccttagacta atagacgttt tggaaagttc ggctagtgtt cgtgtgtttg tcgtagcacc    240
cagagcctcc accaaaccct ctccatgtct ttacctcca gtcgctctaa gatctgcttg    300
aagtctcgta tttgtactgc tttctgcttt tctcccaccc ctccctagcac ccccatcc    360
cccatctagt aacatctcag aaatttcac cagaggaaca aaaaaattaa aaatagaaca    420
tagcaaagca aagacagaat gccccccccc aaatatgttc ctgtccctgt ctgggagttg    480
tgttatttaa agatattctg tatgttgtat cttttgcatg tagcttcctt aat        533
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<210> 94

<211> 413

<212> DNA

<213> Homo Sapiens

<400> 94

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atctggaagg ctctgatcca cctgagcgac ctccgggagt acaggcgctt tgagaaggag    60
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33

aagctcaagt ccagtgga caatgataat ccccttttca agagcgccac cagcagggtc 120
 atgaacccca agtttctga gagttaggag cacttggtga agacaaggcc gtcaggaccc 180
 accatgtctg ccccatcacg cggccgagac atggcttggc cacagctctt gaggatgtca 240
 ccaattaacc agaaatccag ttattttccg cctcaaaat gacagccatg gccggccgggt 300
 gcttctgggg gctcgtcggg gggacagctc cactctgact ggacagctct ttgcatggag 360
 acttgaggag ggcttgagggt tggtagggtt aggtgcgtgt ttctctgtgca agt 413

<210> 95
 <211> 465
 <212> DNA
 <213> Homo Sapiens

<400> 95
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 ccacccgagg gccagtgagt cggggaagag gacttctcac tcccagagca agaggagtcc 120
 cccaactgg gtacagacct ccaccgccac ccccgacaca agagacttat ggagaatatg 180
 actatgatga tggatatggc actgcttatg atgaacagag ttatgattcc tatgataaca 240
 gctatagcac ccagcccaa agtgggtctg attactatga ttacggacat ggactcagtg 300
 aggagactta tgattctcac gggcaagaag agtggactaa ctcaagacac aaggcacctt 360
 cagcgaggac agcaaagggc gtctacagag accagccata tggcagatac tgattgtact 420
 gtctgatgtt gtgaaatagc caatctccac cagtctgtga tactg 465

<210> 96
 <211> 537
 <212> DNA
 <213> Homo Sapiens

<400> 96
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 ctcatcagca gcatcgaggc ccagctgagc gagctccgca gtgagatgga gtgccagaac 120
 caagagtaca agatgctgct ggacatcaag acacgtctgg agcaggagat cgccacctac 180
 cgcagcctgc tcgagggcca ggacgccaa gacgctcagc ccccgtagca cctctgttac 240
 cagcacttct agtgcctctg ttaccaccac ctctaagcc tctggctgcc gcaacttctga 300
 tgtccgtagg ccttaaactc gcctggcgtc ccctccctct gtcttcagca ccagaggag 360
 gagagagccg gcagttccct gcaggagaga ggaggggctg ctggacccaa ggctcagtc 420
 ctctgtctc aggacccctt gtctgactc tctctgatg gtgggccctc tgtgctcttc 480
 tcttccggtc ggatctctct cctctctgac ctggatacgc tttggtttct caacttc 537

<210> 97
 <211> 372
 <212> DNA
 <213> Homo Sapiens

<400> 97
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 caaagtttca taaagccctt aagctcatga ttttcatcaa ctctttgccc acatagtcac 120
 ttacctccac agccgtttgt tgtcatagaa ggggtgggtg tgtttggatt tgattttttt 180

34

caacttgtag tgagaaatag gataggtgac aaaaccttac ttgttttctt aagacaattc 240
 agtgcttgag catctctgtc agaaatggaa tgaaatactg ttagccaatt agaattatct 300
 tatgtattgt tattgtgttt tgctgatttt tatatgaaaa tataattatt cattcttgat 360
 ctctggaagc aa 372

<210> 98
 <211> 365
 <212> DNA
 <213> Homo Sapiens

<400> 98
 gggagccaag gctttatagc tctaaagaaa atattcagta gctgaatccg cccagtgata 60
 gcctgtgggc accagcagca agggctgcca tgggatacag caccatctc caaagacctc 120
 tattacataa aactgcttc ttacaggaaa caaacctctt ctgggatctc cttttgtgaa 180
 aaccagtttg atgtgctaaa agtaaaaagt ctattttcca gtgtggtctt gttcagaagc 240
 agccagattt ccaatgttgt ttttccctc cactcagaaa cccctgccct ttcccttcag 300
 aaaacgatgg caggcattcc tctgagttta caagcagaga ctactccaa cccaaactag 360
 ctggg 365

<210> 99
 <211> 465
 <212> DNA
 <213> Homo Sapiens

<220>
 <221> misc_feature
 <222> (110)..(110)
 <223> n is a, c, g, or t

<400> 99
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 attgatgata gttgaaatat tcttagaagg ggtgtgtatg tctagctgtg tctaccatgt 180
 gtatgtatc ttgacaagca gtataaaata cctgtgattt ttctttacat tagggataat 240
 gcataaggaa ttaatcttca tatatattat catccctaatt gtagcagggg gaagtattta 300
 attgcccatg atatgtatct tactttact atgccagaga ggaaactata aagtaattac 360
 acatgtaatc ttgggttttt cacatatgta ggtattcatt ttgagtaggt tgaagaagaa 420
 aaaaaatatt taaatgaatt gaattcctga tgggatatga tcaat 465

<210> 100
 <211> 515
 <212> DNA
 <213> Homo Sapiens

<400> 100
 gaactctgca tcttcatggt ttacagaaat tgggtgcaggc agccagcagt tagattccat 60
 tcatgtaaca cagttggaga gagataccgt tttagtgtgt ttagacaaat ttgtgaaaat 120
 tgtaaactca caaggaaaaa taaaatcaag taagaaactg gcctctgagt taagttttga 180

35

ttttcgcatt gaatctgtag tatgccttca agacagtgtg ttggctttct ggaaacatgg 240
 gatgcagggt aaaagcttca agtcagatga ggttaccag gagatttcag atgaaacaag 300
 agttttccgc ttattaggat cagacagggt tgtcgttttg gaaagtaggc caacagaaaa 360
 tcctactgca cacagcaatc tctacatctt ggctggacat gaaaatagtt actaagcaac 420
 agaaactgat ctcaaatgac aggaaaatga atatactcca ttgaaaggga aaataaggaa 480
 attcaataca aactgcacta tgatttgcct taact 515

<210> 101
 <211> 525
 <212> DNA
 <213> Homo Sapiens

<400> 101
 ctcagagcca cccctaaaga gatcctttga tattttcaac gcagccctgc tttgggctgc 60
 cctgggtgctg ccacacttca ggctcttctc ctttcacaac cttctgtggc tcacagaacc 120
 cttggagcca atggagactg tctcaagagg gcactgggtg cccgacagcc tggcacaggg 180
 cagtgggaca gggcatggcc aggtggccac tccagacccc tggcttttca ctgctggctg 240
 ccttagaacc tttcttacat tagcagtttg ctttgtatgc actttgtttt tttctttggg 300
 tcttggtttt tttttccact tagaaattgc atttcctgac agaaggactc aggttgtctg 360
 aagtcactgc acagtgcac tcagcccaca tagtgatggg tccccgttc actctactta 420
 gcatgtccct accgagtctc ttctccactg gatggaggaa aaccaagccg tggcttcccg 480
 ctcagccctc cctgcccctc ccttcaacca tccccatgg gaaat 525

<210> 102
 <211> 418
 <212> DNA
 <213> Homo Sapiens

<400> 102
 gcaacaaccg aaaatgcacc agccccaggt cctcggacac cgaggagaat gtcaagagggc 60
 gaacacacaa cgtcttggag gcgcagagga ggaacgagct aaaacggagc ttttttgcce 120
 tgctgaccca gatccccgag ttggaaaaca atgaaaaggc cccaaggta gttatcctta 180
 aaaaagccac agcatacatc ctgtccgtcc aagcagagga gaaaagctc atttctgaag 240
 aggacttggt gcggaaacga cgagaacagt tgaaacacaa acttgaacag ctacggaact 300
 cttgtgcgta aggaaaagta aggaaaacga ttccttctaa cagaaatgtc ctgagcaatc 360
 acctatgaac ttgtttcaaa tgcattgatca aatgcaacct cacaaccttg gctgagtc 418

<210> 103
 <211> 462
 <212> DNA
 <213> Homo Sapiens

<400> 103
 aacatccgcc tggtaccag tcgtcttggc tgggcacttc caccgcacc tcattcctac 60
 atcaatgagt ggctccaaat agacctgggg gaggagaaga tcgtgagggg catcatcatt 120
 cagggtggga agcaccgaga gaacaagggt tcatgagga agttcaagat cgggtacagc 180
 aacaacggct cggactggaa gatgatcatg gatgacagca aacgcaaggc gaagtctttt 240

36

gagggcaaca acaactatga tacacctgag ctgcggaactt ttccagctct ctccacgca 300
ttcatcagga tctaccccga gagagccact catggcggac tggggctcag aatggagctg 360
ctgggctgtg aagtggaaagc ccctacagct ggaccgacca ctccaacgg gaacttggtg 420
gatgaatgtg atgacgacca ggccaactgc cacagtggaa ca 462

<210> 104
<211> 370
<212> DNA
<213> Homo Sapiens

<220>
<221> misc_feature
<222> (168)..(168)
<223> n is a, c, g, or t

<400> 104
gcaaatatct taccaggcag cctatgaatt aacccaaaga agctttgggtt ggttttggtg 60
gatttttatc atgccatggtt ggacatgaga ttttttagat cttccttccc acattgctag 120
acgtctcact caaagacatt tgttgggagt cacatttgca tcataganga gacagtccat 180
tcacttagt taaattggat tgagaatgcc ttttgttcc aggaaaatat tgatcacat 240
gaaagaagaa tagttttttg tccccagaga cattcattta gttgatataa tctaccaga 300
aggaaagcac taagaaacac tcgtttgttg tttttaaagg caacagactt aaagttgtcc 360
tcagccaagg 370

<210> 105
<211> 434
<212> DNA
<213> Homo Sapiens

<400> 105
caggtgtatc tgcacagtgg tcgccccaca gcagaccatg tgttcacggg atgcccgcac 60
aaaacagctg aggcagctac tggagaaggt gcagaacatg tctcaatcca tagaggctt 120
ggacaggcgg acccagagag acttcagta cgtggagaag atggagaacc aaatgaaagg 180
actggagtcc aagttaaagc aggtggagga gaggcataag caacacctgg ccaggcagtt 240
taagggctaa cttaaaagag ttttttcaat gctgcagtga ctgaagaagc agtccactcc 300
catgtaacca tgaaagagag ccagagagct ttttgacca tgcattttta ctattatatt 360
ccaatactta gcaccatttc actaaggaac cttgaatata accaggatcc tcctttgcat 420
gcgactgtag ctgc 434

<210> 106
<211> 503
<212> DNA
<213> Homo Sapiens

<220>
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<222> (158)..(158)
<223> n is a, c, g, or t

<220>

<221> misc_feature
 <222> (216)..(217)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (231)..(231)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (250)..(250)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (261)..(261)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (291)..(291)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (297)..(297)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (341)..(341)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (352)..(352)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (365)..(365)
 <223> n is a, c, g, or t

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 ccccggtttt cctccccgcc ctgtccctct ctggtcaaac aacatactaa agaggcgagg 120
 caatgactgt tggccagttc tcaccgggga aaaacccnac tgtaggatg gcatgaacat 180
 ttccttagat cgtggtcagc tccgaggaat gtggcncca ggctctttga ngagccatgg 240
 gctgcacccn ggccgtagge ntagtgtaac tcgcatccca ttgcagtgcc ngtttctttg 300
 actgtgttgc tgtctcttag attaacctg ctgaggctcc nacatagctc cntggacctg 360
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 ttgtttgaac tcgcgtaccc cgtagataca ttgtgcaacg ttcttctgtt attcccttga 480
 ggtggttaact tcgtatgttc agt 503

<210> 107
 <211> 556
 <212> DNA
 <213> Homo Sapiens

<400> 107
 ggagacttga gcttgacctt aggatatgca ttaaccactc tacagaactcc cactcagtag 60

38

tgtacaggggt ggctgtggtc ctagaagttc agtttttact gaggaatat ttccattaac 120
 agcaattatt atattgaagg ctttaataaa ggccacagga gacattacta tagcatagat 180
 tgtcaaagt aaatttactg agcgtgtttt ataaaaaact cacagggtgt tgaggccaaa 240
 acagatttta gacttacctt gaacggataa gaatctatag ttcactgaca cagtaaaatt 300
 aactctgtgg gtggggggcg ggggcatagc tctaactctaa tatataaaat gtgtgatgaa 360
 tcaacaagat ttccacaatt cttctgtcaa gcttactaca gtgaaagaat gggattggca 420
 agtaacttct gacttactgt cagttgtact tctgtccat agacatcagt attctgcat 480
 catttttgat gactacctca gaacataaaa aggaacgtat atcacataat tccagtcaca 540
 gtttttggtt cctctt 556

<210> 108
 <211> 543
 <212> DNA
 <213> Homo Sapiens

<400> 108
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 acaatttaga agtttgatta attatattat ctatttaggc attaatataa aagaggtagg 180
 agtctgttat ttaaaaaaag cattaaattt aaaaaaaaac tgtcttgtct acttttagct 240
 tcattctccc atattttgaa ggggtgtgtaa cttcagctct gcaggattgc atggggtaaa 300
 acttgttacc aacacatgtg aaccattgct acattgtagg ttgtgatcat tttgccccac 360
 tgaagcccat gtatctgacc ttacgtgcct tttgaactag gagaatcggg ctaatttatt 420
 aatgatgata attataatgt atctgtacag cactttttac atttggaag tgctttccaa 480
 tccatgtagg ttactagtta ttacagctgt aaggataaaa cacgtcatgt ggattcattt 540
 tga 543

<210> 109
 <211> 458
 <212> DNA
 <213> Homo Sapiens

<400> 109
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 gaaacactat ttgtgacttt ttaaacgatt agtgatgtcc ttaaaatgtg gtctgccaat 120
 ctgtacaaaa tggctctatt tttgtgaaga gggacataag ataaatgat gttatacatc 180
 aatatgtata tatgtatttc tatatagact tggagaatac tgccaaaaca tttatgacaa 240
 gctgtatcac tgccttcgtt tatatttttt taactgtgat aatccccaca ggcacattaa 300
 ctgttgcaact tttgaatgtc caaaatttat attttagaaa taataaaaag aaagatactt 360
 acatgttccc aaaacaatgg tgtggtgaat gtgtgagaaa aactaacttg atagggtcta 420
 ccaatacaaa atgtattacg aatgcccctg ttcatggt 458

<210> 110
 <211> 412
 <212> DNA

39

<213> Homo Sapiens

<400> 110

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gtcaaaccat gactcgaca tggcaaaaga acggggccac agtacagcct cacattcttc      60
ttccaattct gaagatacag agatgtgatg aaaacaagta atagctttgg ctgtttatatt    120
gatagctggt tctgggtatt taataggaat cctttctcaa ggaatgagtt gtgacctgtt    180
tactgtctct ttagaagaaa aactccactg gaaaccattc accatgtgtg actgtcttct    240
gttatcattt gtcttacagg cggctattgc agacggctaa tttatgctta acttaggaag    300
agataaggca agagctagat ttttttcatg tgatcttttc caagcttcaa cttaacttaa    360
ctacatttct ctgtatgatg atgtctctta cttctacagg ttccttgagc ac            412

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<210> 111

<211> 514

<212> DNA

<213> Homo Sapiens

<400> 111

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taaattcaca tgcagtctca gagactatth agacaaagtt caagtttagga gcttttagga      60
tgtggggagta aaactttaat gggaggggag ggctggctgc tggagaagag aagaagccag    120
actggttaga cagtactctt aactcctagc ccagcctacg tgccctgccc ctctggccac    180
tgctgcagac acctgcctta acacacacac ctctaggact ccacagtttt gccttaaagg    240
accttcccaa gtctcccttt ccctgtctgg cttctccctt aagaagagag agatacttgt    300
agaattgggt ggggggaatg agcatgaact gtccttccat ttgggatatg ttacattaga    360
gtgagagaga gaataaggag cctttcttat ggaagaaatg ggagaagaga gacagggttc    420
ttttcagcag agtctagtag tttctctgta aggcaaaata atctaaaaag actaacctgc    480
ccaccactc cttatattgc tgtgagattg cccc                                514

```

<210> 112

<211> 489

<212> DNA

<213> Homo Sapiens

<400> 112

```

cggaccatc caagtcatct gattgaagag catgacagaa acaaaatgta ttcaccaagc      60
atthtaggat ttgacttttt cactaaccag ttgacgagca gtgcatttac aaggcactgc    120
caaacaagat gcccttggga gctgtgaggg aaagaggacc tgcgggctta gatcaatctc    180
aattcctttt catgccctcc tgcattgctg ctgcgtgggt atttgtctcc ttagccatca    240
ggtacagttt aactacaat gtaagctata ggtggagcat cagcagtgag tgaggccatt    300
cttcatcctt aggatgtggc aatgaaatga tggtgcaagt tcctttctct tttgtgaatc    360
tttcccccca tttcctgttt acatgtaacc caacaaaatg caatttctag tgccttctgt    420
ccaatcagtt ctttctctg agtgagacgt acttggtctac agatttctgc cttgttttgc    480
gacattgtc                                489

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<210> 113

<211> 416

<212> DNA

<213> Homo Sapiens

40

<400> 113
gattgggatg gccttagctc ttagccaaac accttcctga caccatgagg gccagcagct 60
tcttgatcgt ggtgggtgtc ctcatcgctg ggacgctggt tctagaggca gctgtcacgg 120
gagttcctgt taaaggtcaa gacactgtca aaggccgtgt tccattcaat ggacaagatc 180
ccgttaaagg acaagtttca gttaaaggtc aagataaagt caaagcgcaa gagccagtca 240
aaggteccagt ctccactaag cctgggtcct gccccattat ctggtaccgg tgcgccatgt 300
tgaatcccc taaccgctgc ttgaaagata ctgactgccc aggaatcaag aagtgtgtgtg 360
aaggctcttg cgggatggcc tgtttcgttc ccagtgaa ggagccggtc ctgtct 416

<210> 114
<211> 502
<212> DNA
<213> Homo Sapiens

<400> 114
cccgaacggg gggcatttgt gaggcccatg gttgagaaat gaataatttc ccaattagga 60
agtgtgaagca gctgagggtc cttagaggag cttagccaat gtgggagcag cggtttgggg 120
agcagagaca ctaacgactt cagggcaggg ctctgatatt ccatgaatgt atcaggaaat 180
atatatgtgt gtgtatgttt gcacacttgt tgtgtgggct gtgagtgtaa gtgtgagtaa 240
gagctggtgt ctgattgtta agtctaaata ttctctaaa ctgtgtggac tgtgatgcca 300
cacagagtgg tctttctgga gaggttatag gtcactcctg gggcctcttg ggtccccac 360
gtgacagtgc ctgggaatgt acttattctg cagcatgacc tgtgaccagc actgtctcag 420
tttcaacttc acatagatgt ccctttcttg gccagttatc ccttctttt agcctagttc 480
atccaatcct cactgggtgg gg 502

<210> 115
<211> 430
<212> DNA
<213> Homo Sapiens

<400> 115
accacaacga cattgccttg ctgaagatcc gttccaagga gggcagggtg gcgcagccat 60
cccggactat acagaccatc tgccctgccct cgatgtataa cgatccccag tttggcacia 120
gctgtgagat cactggcttt ggaaaagaga attctaccga ctatctctat ccggagcagc 180
tgaagatgac tgttgtgaag ctgatttccc accgggagtg tcagcagccc cactactacg 240
gctctgaagt caccacaaa atgctgtgtg ctgctgacct acagtggaaa acagattcct 300
gccagggaga ctgaggggga cccctcgtct gttccctcca aggccgcatg actttgactg 360
gaattgtgag ctggggccgt ggatgtgccc tgaaggacaa gccaggcgtc tacacgagag 420
tctcacactt 430

<210> 116
<211> 449
<212> DNA
<213> Homo Sapiens

<400> 116
gggttgccat ccaagtgaag gtcttttctt tgaccaaggg ggacagtcag ttttgcaaaa 60

41

ggactctaatacctgttttaa tattgtcttc ctaattggga taatttaatt aacaagattg 120
 actagaagtgaactgcaac actaacttcc cctgctgtg gtgtgacctg agttggtgac 180
 acagggcaca gacccagag cttggctttt gaaacacaac tcagggtttt tgtgaaggtt 240
 ccccgctga gatctttcct cctgggttact gtgaagcctg ttggtttgc gctgtcgttt 300
 ttgaggagg cccatggggg taggagcagt tgaacctggg aacaaacctc acttgagctg 360
 tgcctagaca atgtgaattc ctgtgttgct aacagaagtgcctgtaagc tcctgtgctc 420
 cggagggaag catttcctgg taggctttg 449

<210> 117
 <211> 535
 <212> DNA
 <213> Homo Sapiens

<400> 117
 gctgaaggca gatgtcgtcc caaagacagc tgagaacttc agagccctgt gcaactggtga 60
 gaagggcttc ggctacaaag gctccacctt ccacaggggtg atcccttcc tcatgtgcca 120
 ggcgggcgac ttcaccaacc acaatggcac aggcgggaag tccatctacg gaagccgctt 180
 tcctgacgag aactttacac tgaagcacgt ggggccaggt gtctgtcca tggctaattgc 240
 tggctctaac accaaccggct ccagttctt catctgcacc ataaagacag actggttgga 300
 tggcaagcat gttgtgttcg gtcacgtcaa agagggcatg gacgtcgtga agaaaataga 360
 atctttcggc tctaagagtg ggaggacatc caagaagatt gtcatcacag actgtggcca 420
 gttgagctaa tctgtggcca ggggtgctggc atgggtggcag ctgcaaatgt ccatgcaccc 480
 aggtggccgc gttgggctgt cagccaagggt gcctgaaacg atacgtgtgc ccaact 535

<210> 118
 <211> 484
 <212> DNA
 <213> Homo Sapiens

<400> 118
 ggttgaatgt ttgtccttag gataggccta tgtgctagcc cacaagaat attgtctcat 60
 tagcctgaat gtgccataag actgacctt taaaatgttt tgagggatct gtggatgctt 120
 cgttaatttg ttcagccaca atttattgag aaaatattct gtgtcaagca ctgtgggttt 180
 taatattttt aaatcaaacg ctgattacag ataatagtat ttatataaat aattgaaaaa 240
 aattttcttt tgggaagagg gagaaaatga aataaatatc attaaagata actcaggaga 300
 atcttcttta caattttacg tttagaatgt ttaagggttaa gaaagaaata gtcaatatgc 360
 ttgtataaaa cactgttcac tgtttttttt aaaaaaaaaa cttgatttgt tattaacatt 420
 gatctgctga caaacctgg gaatttgggt tgtgtatgcg aatgtttcag tgcctcagac 480
 aaat 484

<210> 119
 <211> 495
 <212> DNA
 <213> Homo Sapiens

<400> 119

42

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gaacaagcgt cctggggcat ttgctattta cctggagcct tggcatttag acatctttga      60
attccttgat ttaaagaaga acacaggaaa ggaagagcag cgtgccagag atcttttctt      120
tgctcttttg attccggatc tcttcatgaa acgagtggag actaatcagg actgggtctt      180
gatgtgtcca aatgagtgtc ctggtctgga tgaggtttgg ggagagggaat ttgagaaact      240
atatgcaagt tatgagaaac aaggctcgtgt ccgcaaagtt gtaaaagctc agcagctttg      300
gtatgccatc attgagtctc agacggaaac aggcaccccg tatatgctct acaaagattc      360
ctgtaatcga aagagcaacc agcagaacct gggaaccatc aaatgcagca acctgtgcac      420
agaaatagtg gagtacacca gcaaagatga gggtgctgtt tgtaatttgg cttccctggc      480
cctgaatatg tatgt                                     495

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<210> 120
<211> 438
<212> DNA
<213> Homo Sapiens

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<400> 120
gcccttgagg tcgcgagaaa agggccgtaa ccggaggacc cacgccctg agcctcgcg      60
tgagcggggg ccgcgagcgg caacgcactg gtgaccagac tgtccccacg ccgggaacca      120
agcaggagac gacaggcgag agaggagcca gacagaccct gaaaagaagg acgggttggg      180
gccgggcaca ttgggggtca ccggccgatg gagacaccaa ccgacaggcc ctggctgagg      240
gcagctgcg      300
gggcttattt attaacagga taaccttga atgtagcagc cccgggaggg      360
cggcacaggt cggcgagcgg attcagccgg agggaaggga cggggaagcc gagctccaga      420
gcaacgacca gggccgagga ggtgcctgga gtgcccaccc tgggagacag accccacctc      438
cttgggtagt gagcagtg

```

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<210> 121
<211> 447
<212> DNA
<213> Homo Sapiens

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```

<220>
<221> misc_feature
<222> (116)..(116)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (362)..(362)
<223> n is a, c, g, or t

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<400> 121
ggaactacgg ggcttacagg agcttttgtg tgcttggtag aaactatttc tgttccagtc      60
acattgccat cactcttgta ctgcctgccca ccgcgagga ggctggtgac aggccnaaag      120
gccagtggaa gaaacaccct ttcattctcag agtccactgt ggcaactggc acccctcccc      180
agtacagggg tgctgcaggt ggcagagtga atgtcccca tcatgtggcc caactctcct      240
ggcctggcca tctccctccc cagaacaggt gtgcatgggt tattttggag tgtaggtgac      300
ttgtttactc attgaagcag atttctgctt ctttttattt ttataggaat agaggaagaa      360
angtcagatg cgtgccagc tcttcacccc ccaatctctt ggtggggagg ggtgtaccta      420

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aatattttatc atatccttgc ccttgag 447

<210> 122
 <211> 323
 <212> DNA
 <213> Homo Sapiens

<400> 122
 aaattgacca tacaatttca tcctccttca ggggatcaaa aggacggagt ggggggacag 60
 agactcagat gaggacagag tggtttccaa tgtgttcaat agatttagga gcagaaatgc 120
 aaggggctgc atgacctacc aggacagaac tttccccaat tacaggggtga ctcacagccg 180
 cattggtgac tcacttcaat gtgtcatttc cggtgctgt gtgtgagcag tggacacgtg 240
 aggggggggt ggggtgagaga gacaggcagc tcggattcaa ctaccttaga taatatttct 300
 gaaaacctac cagccagagg gta 323

<210> 123
 <211> 499
 <212> DNA
 <213> Homo Sapiens

<400> 123
 gtatcaggct tcaattccat tatgttttaa tgttgtctct gaagatgact tgtgattttt 60
 ttttcttttt tttaaaccat gaagagccgt ttgacagagc atgctctgcg ttgttggttt 120
 caccagcttc tgccctcaca tgcacaggga tttaacaaca aaaatataac tacaacttcc 180
 cttgtagtct cttatataag tagagtcctt ggtactctgc cctcctgtca gtagtggcag 240
 gatctattgg catattcggg agcttcttag agggatgagg ttctttgaac acagtgaaaa 300
 tttaaattag taactttttt gcaagcagtt tattgactgt tattgctaag aagaagtaag 360
 aaagaaaaag cctgttgcca atcttggtta tttctttaag atttctggca gtgtgggatg 420
 gatgaatgaa gtggaatgtg aactttgggc aagttaaattg ggacagcctt ccatgttcat 480
 ttgtctacct cttaactga 499

<210> 124
 <211> 328
 <212> DNA
 <213> Homo Sapiens

<400> 124
 taattttaga ttcgccttac aatgtaaatc ttcacattgg agataatatt ggttggacct 60
 tgcccatctt cactctagcc ttcgtatttg tgaaggactc agccaccttc cttcttcacc 120
 ccatgcttct caccaaattt ttgttgcac tgagggcact tggataactc aagttgatat 180
 ttatagctga tcaatctata tgtgtcacag aactatgctg cctaaagtga tcttggctcc 240
 ttaatggctc ttttggcccc ttggatagtt aacagctgag taattctaatt ctcttctgtg 300
 ttttccttgc cttaaccaca aattgtgg 328

<210> 125
 <211> 489
 <212> DNA
 <213> Homo Sapiens

44

<400> 125
gagatacaga acttggtgac ccatgtattg cataagctaa agcaacacag acactcctag 60
gcaaagtttt tgtttgtgaa tagtacttgc aaaacttgta aattagcaga tgactttttt 120
ccattgtttt ctccagagag aatgtgctat atttttgtat atacaataat atttgcaact 180
gtgaaaaaca agttgtgcca tactacatgg cacagacaca aaatattata ctaatatgtt 240
gtacattcgg aagaatgtga atcaatcagt atgttttttag attgtatttt gccttacaga 300
aagcctttat tgtaagactc tgatttcctt ttggacttca tgtatattgt acagttacag 360
taaaattcaa cctttatttt ctaatttttt caacatattg tttagtgtaa agaattttta 420
tttgaagttt tattatttta taaaaagaa tattttattt aagaggcatc ttacaaattt 480
tgccccctt 489

<210> 126
<211> 503
<212> DNA
<213> Homo Sapiens

<400> 126
gcggcatgtg accatcattg aactggtggg acagccacct caggagggtg ggcgcatccg 60
ggagcaacag ctgtcagcca acatcatcga ggagctcagg caatttcagc gcctcactcg 120
ctcctaacttc aacatggtgt tgattgacaa gcagggtatt gaccgagacc gctacatgga 180
acctgtcacc cccgaggaaa tcttcacatt cattgatgac tacctaactga gcaatcagga 240
gttgaccag cgtcgggagc aaagggacat atgcgagtga acttgagcca gggcatggtt 300
aaagtcaagg gaaaagctcc tctagttagc tgaaactggg acctaataaa aggaggaaat 360
gttttccac agttctaggg acaggactct gagtggttg agtttgacaa atcctgcagt 420
gtttccaggc atccttttag gactgtgtaa tagtttcctt agaagctagg tagggactga 480
ggacaggcct tgggcagtgg gtt 503

<210> 127
<211> 436
<212> DNA
<213> Homo Sapiens

<400> 127
agactgggag aaaggctgtc cggagggcag accagggtgcc ttgccgcaga gaaaacacca 60
aagtctcctg ttcgctcata aagaagtttt tgggatggga gagaatccag accatcttgg 120
ggcagccagg cccttgccct cattttttaca gaggtagcac aactgattcc aacacaaaac 180
cccttccctt ttttaaatg atttctgttc taatgccata gatcaaaggc ctcaaaaacc 240
attgtgtgtt tcctctttga agcaatgaca agcactttac tttcacgggtg gtttttgttt 300
tttcttattg ctgtggaacc tcttttgag gacgttaaag gcgtgtttta cttgtttttt 360
taagagtgtg tgatgtgtgt tttgtagatt tcttgacagt gctgtaatac agacggcaat 420
gcaatagcct atttaa 436

<210> 128
<211> 497
<212> DNA
<213> Homo Sapiens

<400> 128
 cctgccctct agttggttct gggctttgat ctcttccaac ctgccagtc acagaaggag 60
 gaatgactca aatgccccaa accaagaaca cattgcagaa gtaagacaaa catgtatatt 120
 tttaaatggt ctaacataag acctgttctc tctagccatt gatttaccag gctttctgaa 180
 agatctagtg gttcacacag agagagagag agtactgaaa aagcaactcc tcttcttagt 240
 cttaataatt tactaaaatg gtcaactttt cattatcttt attataataa acctgatgct 300
 tttttttaga actccttact ctgatgtctg tatatgttgc actgaaaagg ttaatatatta 360
 atgttttaat ttattttgtg tggtaagtta attttgattt ctgtaatgtg ttaatgtgat 420
 tagcagttat tttccttaat atctgaatta tacttaaaga gtagtgagca atataagacg 480
 caatgtgtt tttcagt 497

<210> 129
 <211> 321
 <212> DNA
 <213> Homo Sapiens

<400> 129
 gtttggtggtg tggaaggctc cattttattg agatttttaa gatacatgca aagggttgga 60
 aatagaacct ctaggcaccc tctcagtggt ggggtggctg agagttaaag acagtgtggc 120
 tgcagtagca tagaggcgcc tagaaattcc acttgaccg tagggcatgc tgataccatc 180
 ccaatagctg ttgccattg acctctagtg gtgagtttct agaatactgg tccattcatg 240
 agatattcaa gattcaagag tattctcact tctgggttat cagcataaac tggaatgtag 300
 tgtcagagga tactgtggct t 321

<210> 130
 <211> 553
 <212> DNA
 <213> Homo Sapiens

<400> 130
 tttgcctgca gtttcttgtg tagatttgaa aattgtatac caatgtgttt tctgtagact 60
 ctaagataca ctgcactttg tttagaaaaa aaactgaaga tgaaatatat attgtaaaga 120
 agggatatta agaactttag ataacttctt gaaaaagatg gcttatgtca tcagtaaagt 180
 acctttatgt tatgaggata taatgtgtgc tttattgaat tagaaaatta gtgaccatta 240
 ttcacagggtg gacaaatggt gtctctgttaa tttataggag ttttttgggg atgtggaggt 300
 agttgggtag aaaaattatt agaacattca cttttgttaa cagtatttct cttttattct 360
 gttatatagt ggatgatata cacagtggca aaacaaaagt acattgctta aaatatatag 420
 tgaaaaatgt cactatatct tcccatttaa cattgttttt gtatattggg ttagatttc 480
 tgacatcaaa acttggaccc ttggaaaaca aaagttttta ttaaaaaaa tccttgtgac 540
 ttacaatttg cac 553

<210> 131
 <211> 419
 <212> DNA
 <213> Homo Sapiens

46

<400> 131
gagtcggaga tgatgcagca cacacacaat tccccagccc agtgatgctt gtgttgacca 60
gatgttcctg agtctggagc aagcaccag gccagaataa cagagcttcc ttagttgggtg 120
aagacttaaa catctgcctg aggtcaggag gcaatttgcc tgccttgtag aaaagctcag 180
gtgaaagact gagatgaatg tctttcctct ccctgcctcc caccagactt cctcctggaa 240
aacgctttgg tagatttggc caggagcttt cttttatgta aattggataa atacacacac 300
catacactat ccacagatat agccaagtag atttgggtag aggatactat ttccagaata 360
gtgttttagct cacctagggg gatatgtttg tatacacatt tgcataatac cacatgggg 419

<210> 132
<211> 414
<212> DNA
<213> Homo Sapiens

<400> 132
ttgttgtcgt tgcttgtttg aagaaaatca tgacattcca agttgacatt ttttttttca 60
ttttaattaa aatttgaat tctgaacacc gtcagcacc tctcttccct atcatgggtc 120
atctgacccc tgtccgtctc cttgtccctg cttcatgttt gggggccttt ctttaactgc 180
cttctgggt tagctcagat ggcagatgag agtgtagtca agggcctggg cacaggaggg 240
agagctgcag agtgtcctgc ctgccttggc tggagggaca cctctcctgg gtgtggagac 300
agcttgggtc cctttcccta gctccctggt ggggtgaatgc cacctcctga gacccacc 360
tcttgaatt aaaattgttg gtcactgggg aaagcctgag tttgcaacca gttg 414

<210> 133
<211> 419
<212> DNA
<213> Homo Sapiens

<400> 133
aggggtgaa ctatcggtat cacctgggtt gtaactgcaa gatcaagtcc tgctactacc 60
tgccttgctt tgtgacttcc aagaacgagt gtctctggac cgacatgctc tccaatttcg 120
gttaccctgg ctaccagtcc aaacactacg cctgcatccg gcagaagggc ggctactgca 180
gctggtaccg aggatggggc cccccgata aaagcatcat caatgccaca gaccctgag 240
cgccagaccc tgcccacct cacttccctc ccttcccgt gagcttccct tggacactaa 300
ctcttcccag atgatgacaa tgaaattagt gcctgttttc ttgcaaattt agcacttgga 360
acatttaaag aaaggctctat gctgtcatat ggggtttatt gggaactatc ctcctggcc 419

<210> 134
<211> 493
<212> DNA
<213> Homo Sapiens

<400> 134
gacttttttg aatagccctg tctagggcaa actgtggccc ccaggagaca ctacccttcc 60
atgccccaga cctctgtctt gcatgtgaca attgacaatc tggactaccc caagatggca 120
cccaagtgtt tggcttctgg ctacctaagg ttaacatgtc actagagtat ttttatgaga 180
gacaaacatt ataaaaatct gatggcaaaa gcaaaacaaa atggaagta ggggaggtgg 240

47

atgtgacaac aacttccaaa ttggctcttt ggaggcgaga ggaaggggag aacttggaga 300
 atagtttttg ctttgggggt agaggcttct tagattctcc cagcatccgc ctttcccttt 360
 agccagtctg ctgtcctgaa acccagaagt gatggagaga aaccaacaag agatctcgaa 420
 ccctgtctag aaggaatgta tttgttgcta aatttcgtag cactgtttac agttttcttc 480
 catgttattt atg 493

<210> 135
 <211> 567
 <212> DNA
 <213> Homo Sapiens

<400> 135
 gagtattact agagctttgc cacctctcca tttttgcctt ggtgctcacc ttaatggcct 60
 aatgcacccc caaacatgga aatataccca aaaaatactt aatagtccac caaaaggcaa 120
 gactgccctt agaaattcta gcctggtttg gagatactaa ctgctctcag agaaagtagc 180
 tttgtgacat gtcataaacc catgtttgca atcaaagatg ataaaataga ttcttatttt 240
 tccccacccc ccgaaaatgt tcaataatgt cccatgtaaa acctgctaca aatggcagct 300
 tatacatagc aatggtaaaa tcatcatctg gatttaggaa ttgctcttgt catacccca 360
 agtttctaag atttaagatt ctccctacta ctatcctacg tttaaatac tttgaaagtt 420
 tgtattaaat gtgaatttta agaaataata tttatatattc tgtaaatgta aactgtgaag 480
 atagtataaa actgaagcag atacctggaa ccacctaaag aacttccatt tatggaggat 540
 ttttttgcct cttgtgtttg gaattat 567

<210> 136
 <211> 479
 <212> DNA
 <213> Homo Sapiens

<220>
 <221> misc_feature
 <222> (441)..(441)
 <223> n is a, c, g, or t

<400> 136
 accaagggtc tcatgaatct ccaaccttaa atcctgaaac agtggcaata aatttatctg 60
 atgttgactt gagtaaata atcaccacta ttgctggagt catgacacta agtcaagtta 120
 aaggctttgt tcgaaagaat ggtgtcaatg aagccaaaat agatgagatc aagaatgaca 180
 atgtccaaga cacagcagaa cagaaagttc aactgcttcg taattggcat caacttcatg 240
 gaaagaaaaga agcgtatgac acattgatta aagatctcaa aaaagccaat ctttgtactc 300
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48

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<211> 248
<212> DNA
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cgcaacaatc catctctcaa gtagtgtatc acagtagtag cctccagggt tccttaaggg 180
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<213> Homo Sapiens

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ttctcaaact taaatgtcct ctgggaatcc agacttaaaa ataagagcaa acttaacaca     180
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ccagctttcc atcaaactct caatccttga atccaggtaa aagggttaatt atcctaggat     300
tagtgaatga ttcaatgaag ctttcttgaa aacaaacata ggagtgtaat gtactattat     360
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<210> 141

<211> 518

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<213> Homo Sapiens

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ggttcttcta aatgaagcag caacttcgaa aggcgatgtt ggaaaaagac ggataatttg     360
cctagtggga ttgggcctgg tggcttctct cttcagtttt ctactttcaa tatttcgttc     420
caagtaccac ggctatcctt atagtgatct ggactttgag tgagaagatg tgatttggac     480
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<211> 443

<212> DNA

<213> Homo Sapiens

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agaatcacac attattctgg tactgaatgc cgctatcacc atggggatgg ttcttctaaa     180
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